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- ΤI COMPARATIVE IMMUNOCHEMICAL STUDY OF SOME HUMAN LEUKOCYTE INTERFERONS.
- ANSWER 136 OF 142 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on L3 DUPLICATE 47
- LIMITED PROTEOLYSIS OF HUMAN LEUKOCYTE INTERFERON TΤ ALPHA-2 AND LOCALIZATION OF THE MONOCLONAL ANTIBODY BINDING ANTIGENIC DETERMINANT.
- L3ANSWER 137 OF 142 CAPLUS COPYRIGHT 2007 ACS on STN
- Secretion of foreign proteins from Saccharomyces cerevisiae directed by TI α -factor gene fusions
- ANSWER 138 OF 142 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on L3STN **DUPLICATE 48**

- TI BIOLOGIC ACTIVITY IN A FRAGMENT OF RECOMBINANT HUMAN INTERFERON ALPHA.
- L3 ANSWER 139 OF 142 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Mouse interferon receptors: a difference in their response to α and β interferons
- L3 ANSWER 140 OF 142 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 49
- TI Interferon-mediated inhibition of production of Gazdar murine sarcoma virus, a retrovirus lacking env proteins and containing an uncleaved gag precursor
- L3 ANSWER 141 OF 142 CAPLUS COPYRIGHT 2007 ACS on STN
- TI A new mass-spectrometric C-terminal sequencing technique finds a similarity between γ -interferon and α 2-interferon and identifies a proteolytically clipped γ -interferon that retains full antiviral activity
- L3 ANSWER 142 OF 142 CAPLUS COPYRIGHT 2007 ACS on STN
- TI The kidney is the main site of interferon degradation
- => Log off H
 SESSION WILL BE HELD FOR 120 MINUTES
 STN INTERNATIONAL SESSION SUSPENDED AT 13:42:49 ON 25 MAR 2007

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID: SSPTAEGS1646

PASSWORD:

* * * * * RECONNECTED TO STN INTERNATIONAL * * * * * * SESSION RESUMED IN FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE' AT 13:56:52 ON 25 MAR 2007

FILE 'BIOSIS' ENTERED AT 13:56:52 ON 25 MAR 2007

Copyright (c) 2007 The Thomson Corporation

FILE 'CAPLUS' ENTERED AT 13:56:52 ON 25 MAR 2007

COPYRIGHT (C) 2007 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'EMBASE' ENTERED AT 13:56:52 ON 25 MAR 2007

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FILE 'MEDLINE' ENTERED AT 13:56:52 ON 25 MAR 2007

COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 50.90 51.11

FULL ESTIMATED COST

=> D hist

(FILE 'HOME' ENTERED AT 13:34:30 ON 25 MAR 2007)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE' ENTERED AT 13:34:48 ON 25 MAR 2007 L1 70256 S (INTERFERON ALPHA) OR IFN-ALPHA AND (MUTEIN OR VARIANT OR MUT

L2 248 S L1 AND PROTEOL?

L3 142 DUP REM L2 (106 DUPLICATES REMOVED)

=> S L3 and Resistance

L4 · 21 L3 AND RESISTANCE

=> D Ti L4 1-24

- L4 ANSWER 1 OF 21 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
- TI Adenoviral-mediated interferon a overcomes resistance to the interferon protein in various cancer types and has marked bystander effects.
- L4 ANSWER 2 OF 21 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
- TI Gann Monograph on Cancer Research: SPECIAL ISSUE IN COMMEMORATION OF THE 100TH ANNIVERSARY OF THE LATE DR. TOMIZO YOSHIDA'S BIRTH.
- L4 ANSWER 3 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Use of gene expression data and other biochemical criteria in predicting responsiveness to chemotherapy in breast cancer patients
- L4 ANSWER 4 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Genes showing altered levels of expression in pancreatic disease and their use in diagnosis and prognosis of pancreatic cancer
- L4 ANSWER 5 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Truncated polypeptide N-acetylgalactosaminyltransferase II polypeptides and nucleic acids
- L4 ANSWER 6 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Truncated sialyltransferase ST6GalNAc I polypeptides and nucleic acids
- L4 ANSWER 7 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Differentially expressed gene profile for diagnosing and treating mental disorders
- L4 ANSWER 8 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Sequences of human schizophrenia related genes and use for diagnosis, prognosis and therapy
- L4 ANSWER 9 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Gene expression profiles and biomarkers for the detection of Chagas disease and other disease-related gene transcripts in blood
- L4 ANSWER 10 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Gene expression profile of human and mouse genes in atopic dermatitis and psoriasis patients and its use for diagnosis, therapy, and drug screening
- L4 ANSWER 11 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI High throughput directed evolution of proteins and peptides using two-dimensional rational mutagenesis scanning
- L4 ANSWER 12 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Ligand binding domains of cytokine which are linked via flexible polypeptide linker and uses in therapy
- L4 ANSWER 13 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Endocrine disruptor screening using DNA chips of endocrine disruptor-responsive genes
- L4 ANSWER 14 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Long-acting cytokine derivatives and their pharmaceutical compositions
- L4 ANSWER 15 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Gene probes used for genetic profiling in healthcare screening and planning
- L4 ANSWER 16 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Gene probes used for genetic profiling in healthcare screening and planning
- L4 ANSWER 17 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Therapeutic intervention with complement and β -glucan in cancer

- L4 ANSWER 18 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Cloning and cDNA sequences of human interferon .alpha $./\beta$ -binding proteins I and II and their pharmaceutical uses
- L4 ANSWER 19 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Interferon-alpha/beta binding protein, its preparation and use
- L4 ANSWER 20 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Resistance of recombinant proteins to proteolysis during folding and in the folded state
- L4 ANSWER 21 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Identification and partial characterization of a novel protease in Saccharomyces cerevisiae which cleaves the peptide bond between residues 22 and 23 in α -interferon, and identification of an α -interferon resistant to said proteolysis

=> D Ibib ABS L4 1-21

L4 ANSWER 1 OF 21 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN

ACCESSION NUMBER: 2007:206271 BIOSIS DOCUMENT NUMBER: PREV200700198033

TITLE: Adenoviral-mediated interferon a overcomes

resistance to the interferon protein in various cancer types and has marked bystander effects.

AUTHOR(S): Zhang, X.; Yang, Z.; Dong, L.; Papageorgiou, A.; McConkey,

D. J.; Benedict, W. F. [Reprint Author]

CORPORATE SOURCE: Univ Texas, MD Anderson Canc Ctr, Dept Genitourinary Med

Oncol, 1515 Holcombe Blvd, Box 1374, Houston, TX 77030 USA

wbenedic@mdanderson.org

SOURCE: Cancer Gene Therapy, (MAR 2007) Vol. 14, No. 3, pp.

241-250.

ISSN: 0929-1903.

DOCUMENT TYPE: LANGUAGE: Article English

ENTRY DATE:

Entered STN: 21 Mar 2007

Last Updated on STN: 21 Mar 2007

We have previously shown that intravesical administration of adenovirus encoding human interferon alpha-2b (Ad-IFN) induced a marked regression of superficial human bladder tumors derived from cells that are resistant to over 1 million units/ml of IFN alpha protein in vitro. In addition, Ad-IFN appeared to produce strong bystander effects. In this study, we show that Ad-IFN causes marked inhibition of cell growth and apoptosis in cells of various tumor types, all of which are resistant to IFN alpha protein. In addition, strong perinuclear IFN staining was seen in all cell lines following Ad-IFN transfection and was never observed after exposure to the IFN protein. Ad-IFN induced proteolytic processing of caspases 3, 8 and 9, indicative of enzymatic activation. However, the caspase-8-selective inhibitor, IETDfmk, blocked apoptosis only in the cell lines that were sensitive to the IFN alpha protein and had minimal effect on Ad-IFN-induced caspase-3 or -9 processing and cell death, indicating that death receptor-independent mechanism(s) were involved in the cytotoxic effects observed for cancer cell lines resistant to the IFN alpha protein. Moreover, we document that a yet to be identified soluble factor(s) is responsible for causing the bystander effect observed following Ad-IFN treatment in IFN protein-resistant cancer cells.

L4 ANSWER 2 OF 21 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN ACCESSION NUMBER: 2006:165130 BIOSIS

DOCUMENT NUMBER: PREV200600160152

TITLE: Gann Monograph on Cancer Research: SPECIAL ISSUE IN

COMMEMORATION OF THE 100TH ANNIVERSARY OF THE LATE DR.

TOMIZO YOSHIDA'S BIRTH.

AUTHOR (S):

Tsuruo, T [Editor]; Kitagawa, T [Editor]
Tsuruo, T [Editor]; Kitagawa, T [Editor]. Gann Monograph on

SOURCE:

Cancer Research, (2004) Gann Monograph on Cancer Research: SPECIAL ISSUE IN COMMEMORATION OF THE 100TH

ANNIVERSARY OF THE LATE DR. TOMIZO YOSHIDA'S BIRTH.

Publisher: JAPAN SCIENTIFIC SOC PRESS, 2-10 HONGO, 6-CHOME, BUNKYO-KU, TOKYO, 113, JAPAN. Series: GANN MONOGRAPH ON

CANCER RESEARCH.

ISSN: 0072-0151. ISBN: 3-8055-7816-4(H).

DOCUMENT TYPE: LANGUAGE:

Book English

ENTRY DATE:

Entered STN: 9 Mar 2006

Last Updated on STN: 9 Mar 2006

This 280-page book, entitled 'Cancer Research Front of Japan, 2003' is volume 52 of the Gann Monograph on Cancer Research Series and is a special issue published in commemoration of the late Dr. Tomizo Yoshida's birth, who initiated the first publication of this series in 1966. This volume is structured into 4 major sections and contains 18 individually-authored papers. The focus of the first section is pathology and there are 4 papers in this section that individually discuss: the isolation of p53-target genes and their functional analysis; cell adhesion system and human cancer morphogenesis; gastrointestinal stromal tumor as a model for molecular-based diagnosis and treatment of solid tumors; and stem cells and gastric cancer and the role of gastric and intestinal mixed intestinal metaplasia. Carcinogenesis is the theme of the second section, which contains 4 more specific papers. Topics covered in these 4 papers include: renal carcinogenesis in terms of genotype, phenotype and dramatype; heterocyclic amines as mutagens/carcinogens produced during the cooking of meat and fish; a medium-term rat liver bioassay for rapid in vivo detection of the carcinogenic potential of chemicals; and the metabolic activation of polycyclic aromatic hydrocarbons to carcinogens by cytochromes P450 1A1 and 1B1. Cell biology is the focus of the third section, which contains 6 papers on the topic. These 6 papers individually discuss: NK4 in cancer biology and therapeutics; new aspects of interferon-alpha/beta (IFN-alpha/beta) signaling in immunity, oncogenesis and bone metabolism; tumor formation by genetic mutations of beta-catenin, APC, and axin in the Wnt signaling pathway; regulation of transforming growth factor-beta (TGF-beta) signaling and its roles in tumor progression; vascular endothelial growth factor (VEGF) receptor-2 and its unique signaling and specific ligand, VEGF-2; and the roles of pericellular proteolysis by membrane type-1 matrix metalloproteinase in cancer invasion and angiogenesis. The final section concentrates on chemotherapy and the 4 papers in this section individually discuss the antitumor activity of sugar-modified cytosine nucleosides; molecular targeting therapy of cancer in terms of drug resistance , apoptosis and survival signal; the basic and clinical implications of ABC transporters, Y-box-binding protein-1 (YB-1) and angiogenesis-related factors in human malignancies; and molecular mechanisms of angiogenesis in non-small cell lung cancer, and therapeutics targeting related molecules. The book is indexed by author and by subject, and contains 59 figures, 18 of which are in color, and 16 tables. This book will be of interest to oncologists, tumor biology researchers, cell biologists, toxicologists, pathologists and pharmacologists.

ANSWER 3 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2006:795782 CAPLUS

DOCUMENT NUMBER:

145:208138

TITLE:

Use of gene expression data and other biochemical criteria in predicting responsiveness to chemotherapy

in breast cancer patients

INVENTOR(S): PATENT ASSIGNEE(S): Dai, Hongyue; Friend, Stephen H.; Deutsch, Paul Rosetta Inpharmatics LLC, USA; Merck & Co., Inc.

SOURCE:

PCT Int. Appl., 349pp.

CODEN: PIXXD2

DOCUMENT TYPE: LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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PATENT NO.
                KIND
                      DATE
                                APPLICATION NO.
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                               WO 2006-US4280
                A2
                      20060810
WO 2006084272 .
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      SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC,
      VN, YU, ZA, ZM, ZW
   RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
      IS, IT, LT, LU, LV, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ,
      CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG, BW, GH,
      GM, KE, LS, MW, MZ, NA, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
      KG, KZ, MD, RU, TJ, TM
```

PRIORITY APPLN. INFO.:

US 2005-650365P P 20050204

A method of predicting the responsiveness of a breast cancer patient to chemotherapy using a combination of biochem. criteria, especially estrogen receptor levels, age, and gene expression profiles is described. The invention also provides a method for selecting patients for enrollment in a clin. trial of a drug for treating breast cancer based on these factors. Methods of statistical anal. and integration of these data are described.

ANSWER 4 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN L4

ACCESSION NUMBER:

2006:238155 CAPLUS

DOCUMENT NUMBER:

144:310062

TITLE:

Genes showing altered levels of expression in

pancreatic disease and their use in diagnosis and

prognosis of pancreatic cancer

INVENTOR(S):

Kloeppel, Guenter; Luettges, Jutta; Kalthoff, Holger;

Ammerpohl, Ole; Gruetzmann, Robert; Pilarsky, Christian; Saeger, Hans Detlev; Alldinger, Ingo

PATENT ASSIGNEE(S): Technische Universitaet Dresden, Germany

SOURCE:

Ger. Offen., 132 pp.

CODEN: GWXXBX

DOCUMENT TYPE:

Patent German

LANGUAGE:

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATI	ENT :	NO.			KIN	D :	DATE			APPL	ICAT:	ION I	. 00		D	ATE		
	DE :	1020	04042	2822		A1	-	2006	0316		DE 2	004-	1020	0404	2822	20	00408	B31	
	WO 2006024283 A2 WO 2006024283 A3						20060309		1	WO 2005-DE1527						20050826			
	110 2	W:	ΑE,	AG,		AM,	AT,	AU,	AZ,								CA,		
																	GB, KR,		
			LC,	LK,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	
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		RW:	-	-		-		•			•	•	•	•	•	•	HU, BF,	•	
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PRIORITY APPLN. INFO.: DE 2004-102004042822A 20040831 Genes showing altered levels of expression in healthy vs. neoplastic

pancreas are identified for use in the diagnosis of cancers including ductal adenocarcinoma; as indicators in screening for effective drugs; and as targets for nucleic acid-based therapies including antisense nucleic acids or siRNA. Gene expression profiling identified 1419 genes showing changes in levels of expression in neoplastic epithelium of which 650 were up-regulated and 769 were down-regulated. Of the 1419 genes, 1267 were not previously known to have any connection with pancreatic neoplasms.

REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 5 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2005:1330475 CAPLUS

DOCUMENT NUMBER: 144:65957

TITLE: Truncated polypeptide N-acetylgalactosaminyltransferas

e II polypeptides and nucleic acids

INVENTOR(S): Johnson, Karl F.; Chen, Xi; Taudte, Susann; Saribas,

Sami

PATENT ASSIGNEE(S): Neose Technologies, Inc., USA

SOURCE: PCT Int. Appl., 123 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.				KIND		DATE		1	APPL:	ICAT:		DATE					
WO 2005121331				A2		20051222		,	WO 2	005-1		20050603					
WO 2	WO 2005121331				A8		2006	0309									
	W:	ΑE,	AG,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	BG,	BR,	BW,	BY,	BZ,	CA,	CH,
		CN,	CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	EG,	ES,	FI,	GB,	GD,
		GE,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	ΚE,	KG,	KM,	KP,	KR,	KZ,
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		NG,	NI,	NO,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,
		SL,	SM,	SY,	TJ,	TM,	TN,	TR,	TT,	TZ,	UA,	UG,	US,	UZ,	VC,	VN,	YŲ,
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	RW:	BW,	GH,	GM,	KE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,
		ΑZ,	BY,	KG,	ΚZ,	MD,	RU,	ТJ,	TM,	AT,	ΒE,	BG,	CH,	CY,	CZ,	DE,	DK,
		EE,	ES,	FI,	FR,	GB,	GR,	HU,	ΙE,	IS,	ΙT,	LT,	LU,	MC,	ΝL,	PL,	PT,
		RO,	SE,	SI,	SK,	TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GA,	GN,	GQ,	GW,	ML,
		MR,	ΝĖ,	SN,	TD,	TG											
RTTY	APP	IN.	TNFO	. :					1	US 20	004-	5465	30P	1	P 20	0040	603

PRIORITY APPLN. INFO.:

US 2004-546530P P 20040603 US 2004-598584P P 20040803

The present invention features compns. and methods related to mutants of human polypeptide N-acetylgalactosaminyltransferase II (GalNAcT2) that are truncated by deletion of the N-terminal 1-40, 1-73, or 1-94 residues. Truncated forms of GalNAcT2 possess biol. activities comparable to, and in some instances, in excess of their full-length polypeptide counterparts, and may have enhanced properties of solubility, stability, and resistance to proteolytic degradation GalNAcT2 is an essential reagent for glycosylation of therapeutic glycopeptides and oligosaccharides. The invention also features nucleic acids encoding such truncated polypeptides, as well as vectors, host cells, expression systems, and methods of expressing and using such polypeptides.

L4 ANSWER 6 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2005:1330319 CAPLUS

DOCUMENT NUMBER: 144:65956

TITLE: Truncated sialyltransferase ST6GalNAc I polypeptides

and nucleic acids

INVENTOR(S): Johnson, Karl F.; Hakes, David; Wei, Ge; Liu, Li;

Saribas, Sami; Sjoberg, Eric; Clausen, Henrik;

Bennett, Eric Paul; Mobasseri, Aliakbar

PATENT ASSIGNEE(S): . Neose Technologies, Inc., USA

SOURCE: PCT Int. Appl., 192 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent English

LANGUAGE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

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PATENT NO.
                                 KIND
                                           DATE
                                                           APPLICATION NO.
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                                                        WO 2005-US19583
                                           20051222
      WO 2005121332
                                  A2
                                                                                         20050603
            W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
                 CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD,
                 GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KM, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA,
                 NG, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SM, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU,
                 ZA, ZM, ZW
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                 MR, NE, SN, TD, TG
PRIORITY APPLN. INFO.:
                                                           US 2004-576433P
                                                                                     P 20040603
```

US 2005-650011P P 20050204

The present invention features compns. and methods related to mutants of AB human, murine, and chicken CMP-acetylneuraminate- α acetylgalactosaminide $\alpha 2\rightarrow 6$ -sialyltransferase (ST6GalNAcI) that are truncated by deletion of N-terminal residues. Truncated forms of ST6GalNAcI possess biol. activities comparable to, and in some instances, in excess of their full-length polypeptide counterparts, and may have enhanced properties of solubility, stability, and resistance to proteolytic degradation ST6GalNAcI is an essential reagent for glycosylation of therapeutic glycopeptides and oligosaccharides. The invention also features nucleic acids encoding such truncated polypeptides, as well as vectors, host cells, expression systems, and methods of expressing and using such polypeptides.

ANSWER 7 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER:

2005:447673 CAPLUS

DOCUMENT NUMBER:

143:20875

TITLE:

Differentially expressed gene profile for diagnosing

and treating mental disorders

INVENTOR (S):

Akil, Huda; Atz, Mary; Bunney, William E., Jr.; Choudary, Prabhakara V.; Evans, Simon J.; Jones, Edward G.; Li, Jun; Lopez, Juan F.; Myers, Richard; Thompson, Robert C.; Tomita, Hiroaki; Vawter, Marquis

P.; Watson, Stanley

PATENT ASSIGNEE(S):

The Board of Trustees of the Leland Stanford Junior

University, USA

SOURCE:

PCT Int. Appl., 226 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATENT			KIND		DATE		APPLICATION NO.						DATE .				
WO 2005046434					A2 20050526			WO 2004-US36784						20041105			
W :	AE	Ξ,	AG,	AL,	AM,	ΑT,	ΑU,	AZ,	BA,	BB,	BG,	BR,	BW,	BY,	ΒZ,	CA,	CH,
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	GE	Ξ,	GH,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,
	LF	۲,	LR,	LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NA,	NI,
	NC	Ο,	NZ,	OM,	PG,	PH,	PL,	PT,	RO,	RU,	SC,	SD,	SE,	SG,	SK,	SL,	SY,
																ZM,	
RV	V: BV	١,	GH,	GM,	KE,	LS,	MW,	MZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	ΑM,

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AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IS, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
                  NE, SN, TD, TG
       US 2005209181
                                     A1
                                              20050922
                                                               US 2004-982556
                                                                                                 20041104
       AU 2004289247
                                     A1
                                              20050526
                                                               AU 2004-289247
                                                                                                 20041105
       CA 2543811
                                     A1
                                              20050526
                                                               CA 2004-2543811
                                                                                                 20041105
                                                               EP 2004-800741
       EP 1680009
                                     A2
                                              20060719
                                                                                                 20041105
                 AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, PL, SK,
                  HR, IS, YU
PRIORITY APPLN. INFO.:
                                                                US 2003-517751P
                                                                                             P 20031105
                                                               US 2004-982556
                                                                                             Α
                                                                                                 20041104
                                                               WO 2004-US36784
                                                                                             W 20041105
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AB The present invention provides methods for diagnosing mental disorders (e.q., psychotic disorders such as schizophrenia). The present invention uses DNA microarray anal. to demonstrate differential expression of genes in selected regions of post-mortem brains from patients diagnosed with mental disorders in comparison with normal control subjects. The invention also provides methods of identifying modulators of such mental disorders as well as methods of using these modulators to treat patients suffering from such mental disorders.

ANSWER 8 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2005:248643 CAPLUS

DOCUMENT NUMBER: 142:274056

TITLE: Sequences of human schizophrenia related genes and use

for diagnosis, prognosis and therapy

INVENTOR (S): Liew, Choong-Chin

PATENT ASSIGNEE(S): Chondrogene Limited, Can.

SOURCE: U.S. Pat. Appl. Publ., 156 pp., Cont.-in-part of U.S.

Ser. No. 802,875.

CODEN: USXXCO

DOCUMENT TYPE: Patent English LANGUAGE:

FAMILY ACC. NUM. COUNT: 47

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
				-	
US 2004241727	A1	20041202	US 2004-812731		20040330
US 2004014059	A1	20040122	US 2002-268730		20021009
US 2005191637	A1	20050901	US 2004-803737		20040318
US 2005196762	A1	20050908	US 2004-803759		20040318
US 2005196763	A1	20050908	US 2004-803857		20040318
US 2005196764	A1	20050908	US 2004-803858		20040318
US 2005208505	A1	20050922	US 2004-803648		20040318
US 2004241727	A1	20041202	US 2004-812731		20040330
PRIORITY APPLN. INFO.:			US 1999-115125P	P	19990106
			บร์ 2000-477148	B1	20000104
			US 2002-268730	A2	20021009
			US 2003-601518	A2	20030620
			US 2004-802875	A2	20040312
			US 2004-812731	Α	20040330

The present invention is directed to detection and measurement of gene AB transcripts and their equivalent nucleic acid products in blood. Specifically provided is anal. performed on a drop of blood for detecting, diagnosing and monitoring diseases using gene-specific and/or tissue-specific primers. The present invention also describes methods by which delineation of the sequence and/or quantitation of the expression levels of disease-specific genes allows for an immediate and accurate diagnostic/prognostic test for disease or to assess the effect of a particular treatment regimen. [This abstract record is one of 3 records for this document necessitated by the large number of index entries required to fully index the document and publication system constraints.].

L4 ANSWER 9 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2005:139371 CAPLUS

DOCUMENT NUMBER: 142:195820

TITLE: Gene expression profiles and biomarkers for the

detection of Chagas disease and other disease-related

gene transcripts in blood

INVENTOR(S): Liew, Choong-Chin

PATENT ASSIGNEE(S): Chondrogene Limited, Can.

SOURCE: U.S. Pat. Appl. Publ., 154 pp., Cont.-in-part of U.S.

Ser. No. 802,875.

CODEN: USXXCO

DOCUMENT TYPE:

LANGUAGE:

Patent English

FAMILY ACC. NUM. COUNT: 47

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
US 2004241729	A1	20041202	US 2004-813097	•	20040330
US 2004014059	A1	20040122	US 2002-268730		20021009
· US 2005191637	A1	20050901	US 2004-803737		20040318
US 2005196762	A1	20050908	US 2004-803759		20040318
US 2005196763	A1	20050908	US 2004-803857		20040318
US 2005196764	A1	20050908	US 2004-803858		20040318
US 2005208505	A1	20050922	US 2004-803648		20040318
US 2004241729	A1	20041202	US 2004-813097		20040330
PRIORITY APPLN. INFO.:			US 1999-115125P .	P	19990106
			US 2000-477148	B1	20000104
		i	US 2002-268730	A2	20021009
			US 2003-601518	A2	20030620
			US 2004-802875	A2	20040312
			US 2004-813097	Α	20040330

AB The present invention is directed to detection and measurement of gene transcripts and their equivalent nucleic acid products in blood. Specifically provided is anal. performed on a drop of blood for detecting, diagnosing, and monitoring diseases, and in particular Chagas disease, using gene-specific and/or tissue-specific primers. Affymetrix Human Genome U133 and ChondroChip microarrays were used to detect differentially expressed gene transcripts in hypertension, obesity, allergy, systemic steroids, coronary artery disease, diabetes type 2, hyperlipidemia, lung disease, bladder cancer, rheumatoid arthritis, osteoarthritis, liver cancer, schizophrenia, Chagas disease, asthma, and manic depression syndrome. The present invention describes methods by which delineation of the sequence and/or quantitation of the expression levels of disease-specific genes allows for an immediate and accurate diagnostic/prognostic test for disease or to assess the effect of a particular treatment regimen. [This abstract record is one of 3 records for this document necessitated by the large number of index entries required to fully index the document and publication system constraints.].

L4 ANSWER 10 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2004:308529 CAPLUS

DOCUMENT NUMBER:

140:333599

TITLE:

Gene expression profile of human and mouse genes in atopic dermatitis and psoriasis patients and its use

for diagnosis, therapy, and drug screening

INVENTOR(S): Itoh, Mikito; Ogawa, Kaoru; Shinagawa, Akira; Sudo,

Hajime; Ogawa, Hideoki; Ra, Chisei; Mitsuishi, Kouichi

PATENT ASSIGNEE(S): Genox Research, Inc., Japan; Juntendo University

SOURCE:

PCT Int. Appl., 611 pp.

CODEN: PIXXD2

DOCUMENT TYPE: LANGUAGE:

Patent Japanese

FAMILY ACC. NUM. COUNT:

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PATENT NO.
                          KIND
                                  DATE
                                              APPLICATION NO.
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                                            WO 2003-JP9808
     WO 2004031386
                                 20040415
         A1
                                                                       20030801
     AU 2003252326
                           A1
                                  20040423
                                             AU 2003-252326
                                                                       20030801
                                                                   A 20020806
A 20030514
PRIORITY APPLN. INFO.:
                                               JP 2002-229318
                                               JP 2003-136543
                                                                   W 20030801
                                              WO 2003-JP9808
AΒ
     This invention provides gene expression profile between a rash site and a
     no-rash site in a patient with atopic dermatitis or a patient with
     psoriasis. The invention also provides gene expression profile between a
     no-rash site in such a disease and a normal subject. Animal models,
     particularly mouse for those diseases are also claimed. The gene
     expression profile provided in this invention can be used for diagnosis,
     therapy, and drug screening for atopic dermatitis and psoriasis.
REFERENCE COUNT:
                          8
                                 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS
                                 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 11 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER:
                          2004:220463 CAPLUS
DOCUMENT NUMBER:
                          140:265579
TITLE:
                          High throughput directed evolution of proteins and
                          peptides using two-dimensional rational mutagenesis
                          scanning
INVENTOR(S):
                          Gantier, Rene; Guyon, Thierry; Cruz Ramos, Hugo; Vega,
                          Manuel; Drittanti, Lila
PATENT ASSIGNEE(S):
                          Nautilus Biotech, Fr.
                          PCT Int. Appl., 431 pp.
SOURCE:
                          CODEN: PIXXD2
DOCUMENT TYPE:
                          Patent
LANGUAGE:
                          English
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                                 DATE · APPLICATION NO.
     PATENT NO.
                          KIND
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                                           WO 2003-IB4255 20030908
     WO 2004022747
                                20040318
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             TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
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           FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
CA 2498284
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AU 2003267700
                            A1
                                     20040329
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                                                    EP 2003-748392
EP 1539950
                            A1
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                                                  US 2003-658355
US 2005202438
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                                                                                      20030908
                                                    US 2005-196067
US 2006020396
                            A1
                                     20060126
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US 2002-410258P P 20020909

US 2003-457063P P 20030321

US 2002-409898P P 20020909

US 2003-457135P P 20030321
PRIORITY APPLN. INFO.:
                                                 US 2003-658355 A1 20030908
WO 2003-IB4255 W 20030908
                                                 WO 2003-IB4255
AB
     The invention claims processes and systems for the high throughput
     directed evolution of peptides and proteins. It also provides a rational
     method for generating protein variants. The method relies on an indirect
     search for protein improvement for a particular activity, such as
     increased resistance to proteolysis, based on a
     rational amino acid replacement and sequence change at single or a limited
     number of amino acid positions at a time. The target amino acids are
     selected in silico for replacement and are referred to as "is-HIT target
     positions". The collection (or library) of all is-HITs represents the
     first dimension (target residue position) of the two-dimensional scanning
     methods. The second dimension is the replacing amino acids. The
     collection of mutant mols., or in silico candidate LEADS, is generated,
     tested and phenotypically characterized one-by-one, for example in
     addressable arrays. Optimized proteins having modified amino acid
     sequences at some regions along the protein that perform better than the
     starting sequence are identified and isolated. The methods were applied
     to interferon\alpha -2b and interferon-\beta to
     obtain mutants with altered resistance to proteolysis
     and/or higher conformational stability.
REFERENCE COUNT:
                            6
                                  THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS
                                  RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 12 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER:
                            2003:591215 CAPLUS
DOCUMENT NUMBER:
                            139:144956
TITLE:
                            Ligand binding domains of cytokine which are linked
                            via flexible polypeptide linker and uses in therapy
INVENTOR(S):
                           Ross, Richard; Artymiuk, Peter; Sayers, Jon
                         Asterion Limited, UK
PATENT ASSIGNEE(S):
SOURCE:
                           PCT Int. Appl., 37 pp.
                           CODEN: PIXXD2
DOCUMENT TYPE:
                            Patent
LANGUAGE:
                            English
FAMILY ACC. NUM. COUNT: 1
PATENT INFORMATION:
     PATENT NO.
                          KIND
                                   DATE
                                                APPLICATION NO.
                                                                           DATE
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     WO 2003062276
                           A2
     WO 2003062276 A2
WO 2003062276 A3
                                   20030731
                                                WO 2003-GB253
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              GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ,
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          FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                     . A1
                                              CA 2003-2510751
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EP 1468020
                         A2
                                  20041020
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          IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
JP 2005529583 T
                                              JP 2003-562153
                                20051006
                                                                              20030124
IN 2004KN00972
                        Α
                                  20060505
                                                IN 2004-KN972
BR 2004003173 A 20060321 BR 2004-3173
US 2005214762 A1 20050929 US 2005-502344
US 2007054364 A1 20070308 US 2006-595991
                        A1 20050929 US 2005-502344
                                                                             20061113
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PRIORITY APPLN. INFO.:

GB 2002-1679

WO 2003-GB253

W 20030124

US 2005-502344

B3 20050511

AB The invention relates to the provision of oligomeric polypeptides (dimers, trimers, etc) comprising the ligand binding domains of cytokines which are linked via flexible polypeptide linker mols. The linker mols. optionally comprise protease sensitive sites to modulate the release of biol. active cytokines when administered to a human or animal subject. The invention also relates to chemical crosslinkers wherein the chemical crosslinkers serve

link the ligand binding domains. The chimeric cytokine can be used for treating acromegaly, gigantism, GH deficiency, Turners syndrome, renal failure, osteoporosis, diabetes mellitus, cancer, obesity, insulin resistance, hyperlipidemia, hypertension, anemia, autoimmune and infectious disease, inflammatory disorders including rheumatoid arthritis.

L4 ANSWER 13 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2002:937303 CAPLUS

DOCUMENT NUMBER: 138:20443

TITLE: Endocrine disruptor screening using DNA chips of

endocrine disruptor-responsive genes

INVENTOR(S): Kondo, Akihiro; Takeda, Takeshi; Mizutani, Shigetoshi;

Tsujimoto, Yoshimasa; Takashima, Ryokichi; Enoki,

Yuki; Kato, Ikunoshin

PATENT ASSIGNEE(S): Takara Bio Inc., Japan

SOURCE: Jpn. Kokai Tokkyo Koho, 386 pp.

CODEN: JKXXAF

DOCUMENT TYPE: Patent LANGUAGE: Japanese

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE

JP 2002355079 A 20021210 JP 2002-69354 20020313

PRIORITY APPLN. INFO.: JP 2001-73183 A 20010314

JP 2001-74993 A 20010315

JP 2001-102519 A 20010330

AB A method and kit for detecting endocrine-disrupting chems. using DNA

AB A method and kit for detecting endocrine-disrupting chems. using DNA microarrays are claimed. The method comprises preparing a nucleic acid sample containing mRNAs or cDNAs originating in cells, tissues, or organisms which have been brought into contact with a sample containing the endocrine disruptor. The nucleic acid sample is hybridized with DNA microarrays having genes affected by the endocrine disruptor or DNA fragments originating in these genes have been fixed. The results obtained are then compared with the results obtained with the control sample to select the gene affected by the endocrine disruptor. Genes whose expression is altered by tri-Bu tin, 4-octaphenol, 4-nonylphenol, di-N-Bu phthalate, dichlorohexyl phthalate, octachlorostyrene, benzophenone, diethylhexyl phthalate, diethylstilbestrol (DES), and 17- β estradiol (E2), were found in mice by DNA chip anal.

L4 ANSWER 14 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2002:353234 CAPLUS

DOCUMENT NUMBER: 136:359632

TITLE: Long-acting cytokine derivatives and their

pharmaceutical compositions

INVENTOR(S): Shechter, Yoram; Fridkin, Matityahu; Goldwaser, Itzhak

PATENT ASSIGNEE(S): Yeda Research and Development Co., Ltd., Israel

SOURCE: PCT Int. Appl., 43 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

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KIND
       PATENT NO.
                                            DATE
                                                            APPLICATION NO.
                                                                                             DATE
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      WO 2002036067
                                            20020510
                                                            WO 2001-IL1005
                                                                                             20011030
                                   A2
      WO 2002036067
                                   A3
                                            20030109
                 AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
                 GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
                 PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
            RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF,
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                                                          AU 2002-14223
      AU 2002014223
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      US 2004131586
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                                   Α1
                                                                                             20030902
PRIORITY APPLN. INFO.:
                                                             IL 2000-139400
                                                                                        A 20001101
                                                             WO 2001-IL1005
                                                                                        W 20011030
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Cytokine derivs. are provided bearing functional groups sensitive to mild AB basic conditions, such as fluorenylmethoxycarbonyl (Fmoc) and 2-sulfo-9-fluorenylmethoxycarbonyl (FMS), and pharmaceutical compns. comprising them. Preferred derivs, are those in which amino groups of the cytokine are substituted with FMS, for example FMS7-IFN- $\alpha 2$ and FMS3-IL-2. These cytokine derivs. can be administered as inactive or slightly active prodrugs and are capable of undergoing spontaneous regeneration into the parent bioactive drugs under in vivo physiol. conditions and in a homogeneous fashion. The cytokine prodrugs present higher metabolic stability and augmented bioavailability. For example, in an in vivo experiment designed for the evaluation of the anti-metastatic capacity of FMS3-IL-2, mice were inoculated i.v. on day (-3) with 105 D122 metastatic cells. Native IL-2 and FMS3-IL-2 were administered i.p. at high and moderate concns. (5000 ng and 500 ng, resp.) once daily for 30 days. Each group consists of 8 mice. The original protocol for anti-metastatic therapy implies identical dosages given twice a day. However, since prolongation of FMS3-IL-2 in serum is assumed, it is administered only once a day. Metastatic load in lungs of mice was weighed on day 30.

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ANSWER 15 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
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ACCESSION NUMBER: - 1999:795994 CAPLUS

DOCUMENT NUMBER: 132:31744

TITLE: Gene probes used for genetic profiling in healthcare

screening and planning

INVENTOR (S):

Roberts, Gareth Wyn PATENT ASSIGNEE(S): Genostic Pharma Ltd., UK SOURCE: PCT Int. Appl., 745 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

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PATENT NO.
                   KIND
                         DATE
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                                                           DATE
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                                   WO 1999-GB1780
WO 9964627
                   A2
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       JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK,
       MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ,
       TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ,
       MD, RU, TJ, TM
   RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,
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ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
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PRIORITY APPLN. INFO.:
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                                                                  19980819
    There is considerable evidence that significant factor underlying the
AB
     individual variability in response to disease, therapy and prognosis lies
     in a person's genetic make-up. There have been numerous examples relating
     that polymorphisms within a given gene can alter the functionality of the
    protein encoded by that gene thus leading to a variable physiol. response.
     In order to bring about the integration of genomics into medical practice
     and enable design and building of a technol. platform which will enable
     the everyday practice of mol. medicine a way must be invented for the DNA
     sequence data to be aligned with the identification of genes central to
     the induction, development, progression and outcome of disease or physiol.
     states of interest. According to the invention, the number of genes and
     their configurations (mutations and polymorphisms) needed to be identified
     in order to provide critical clin. information concerning individual
     prognosis is considerably less than the 100,000 thought to comprise the
    human genome. The identification of the identity of the core group of
     genes enables the invention of a design for genetic profiling technologies
     which comprises of the identification of the core group of genes and their
     sequence variants required to provide a broad base of clin. prognostic
     information - "genostics". The "Genostic" profiling of patients and
    persons will radically enhance the ability of clinicians, healthcare
    professionals and other parties to plan and manage healthcare provision
     and the targeting of appropriate healthcare resources to those deemed most
              The use of this invention could also lead to a host of new
     applications for such profiling technologies, such as identification of
    persons with particular work or environment related risk, selection of
     applicants for employment, training or specific opportunities or for the
     enhancing of the planning and organization of health services, education
     services and social services.
    ANSWER 16 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
L4
ACCESSION NUMBER:
                         1999:795993 CAPLUS
DOCUMENT NUMBER:
                         132:31743
```

TITLE: Gene probes used for genetic profiling in healthcare

screening and planning

INVENTOR(S):

Roberts, Gareth Wyn

PATENT ASSIGNEE(S): SOURCE:

Genostic Pharma Limited, UK

PCT Int. Appl., 149 pp. CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT:

PATEN'	T NO.			KIN)	DATE			APPL	ICAT	ION 1	NO.		D	ATE	
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                                                                         Α
                                                                            19980606
                                                  GB 1998-28289
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                                                                            19981223
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                                                                            19980805
                                                  GB 1998-17097
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                                                                            1998,0807
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                                                  GB 1998-17200
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                                                  GB 1998-17632
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                                                                            19980814
                                                  GB 1998-17943
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                                                                            19980819
                                                  US ·1999-325123
                                                                         B1 19990603
                                                  WO 1999-GB1779
                                                                         W 19990604
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There is considerable evidence that significant factor underlying the AB individual variability in response to disease, therapy and prognosis lies in a person's genetic make-up. There have been numerous examples relating that polymorphisms within a given gene can alter the functionality of the protein encoded by that gene thus leading to a variable physiol. response. In order to bring about the integration of genomics into medical practice and enable design and building of a technol. platform which will enable the everyday practice of mol. medicine a way must be invented for the DNA sequence data to be aligned with the identification of genes central to the induction, development, progression and outcome of disease or physiol. states of interest. According to the invention, the number of genes and their configurations (mutations and polymorphisms) needed to be identified in order to provide critical clin. information concerning individual prognosis is considerably less than the 100,000 thought to comprise the human genome. The identification of the identity of the core group of genes enables the invention of a design for genetic profiling technologies.

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L4 ANSWER 17 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN
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ACCESSION NUMBER: 1999:360183 CAPLUS

DOCUMENT NUMBER:

131:183492

TITLE:

Therapeutic intervention with complement and

β-glucan in cancer

AUTHOR(S):

Ross, Gordon D.; Vetvicka, Vaclav; Yan, Jun; Xia, Yu;

Vetvickova, Jana

CORPORATE SOURCE:

Department of Microbiology and Immunology, Department of Pathology, Division of Experimental Immunology and Immunopathology, University of Louisville, Louisville,

KY, USA

SOURCE:

Immunopharmacology (1999), 42(1-3), 61-74

CODEN: IMMUDP; ISSN: 0162-3109

PUBLISHER: DOCUMENT TYPE: Elsevier Science B.V.
Journal; General Review

LANGUAGE: English

AB A review and discussion with many refs. Complement (C) has two major effector systems available for host defense. The membrane attack complex

(MAC) generated from components C5-C9 can form membrane-penetrating lesions that lead to cell death by causing a rapid loss of cytoplasmic components. The MAC is only effective against pathogens with outer phospholipid membranes, and cannot kill Gram-pos. bacteria or yeast whose membranes are protected by cell walls. The most important effector mechanism of C is the opsonization of microbial pathogens with the serum protein C3 that leads to their high avidity attachment to the C3-receptors of phagocytic cells. Pathogens that activate complement are first coated with the C3b fragment of C3, which is rapidly proteolyzed into the iC3b fragment by serum factor I. These iC3b fragments serve to promote the high avidity attachment of the 'iC3b-opsonized' pathogens to the iC3b-receptors (CR3, CD11b/CD18) of phagocytic cells and natural killer (NK) cells, stimulating phagocytosis and/or cytotoxic degranulation. Host cells, including neoplastic tumor cells, have been endowed with natural mechanisms for self-protection against both the MAC and the cytotoxic activation of CR3. This review discusses a novel type of immunotherapy for cancer that uses soluble yeast β -glucan to override the normal resistance of iC3b-opsonized tumor cells to the cytotoxic activation of phagocyte and NK cell CR3, allowing this important effector mechanism of the C system to function against tumor cells in the same way that it normally functions against bacteria and yeast. Moreover, the cytotoxic activation of β -glucan-primed NK cell CR3 by iC3b-opsonized tumors is shown to be accompanied by a tumor-localized secretion of the cytokines TNF α , IFN α , IFN γ , and IL-6.

REFERENCE COUNT:

116 THERE ARE 116 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE REFORMAT

L4 ANSWER 18 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1998:667965 CAPLUS

DOCUMENT NUMBER: 129:299458

TITLE: Cloning and cDNA sequences of human interferon

 α / β -binding proteins I and II and

their pharmaceutical uses

INVENTOR(S): Novick, Daniela; Cohen, Batya; Rubinstein, Menachem

PATENT ASSIGNEE(S): Yeda Research and Development Co. Ltd., Israel

SOURCE: U.S., 35 pp., Cont.-in-part of U.S. Ser. No. 115,741,

abandoned.

CODEN: USXXAM

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.		DATE
US 5821078	Α	19981013	US 1995-385191		19950207
US 6458932	B1	20021001	US 1995-472402		19950607
JP 2004254695	A	20040916	JP 2004-90279		20040325
JP 2005162762	A	20050623	JP 2005-4934		20050112
JP 2005200422	Α	20050728	JP 2005-33495		20050209
PRIORITY APPLN. INFO.:			IL 1992-103052	Α	19920903
			IL 1993-106591	Α	19930804
			US 1993-115741	B2	19930903
			IL 1994-108584	Α	19940207
			JP 1993-243987	A3	19930902
			JP 1995-43539	A3	19950207
			US 1995-385191	A3	19950207

AB Interferon α / β binding proteins are provided, which are capable of modulating the activity of interferon- α subtypes as well as interferon- β . Cloning of DNA mols. encoding these proteins, expression in host cells and antibodies against the proteins are also provided. Type I interferons (IFN- α , and IFN- β and IFN- ω) are a family of cytokines usually defined by their ability to confer

resistance to viral infections. There are pathol. situations, related to these cytokines where neutralization of interferon activity may be beneficial to the patient. Cytokine-binding proteins (soluble cytokine receptors) correspond to the extracellular ligand binding domains of their resp. cell surface cytokine receptors. They are derived either by alternative splicing of pre-mRNA common to the cell surface receptor, or by proteolytic cleavage of the cell surface receptor. Therefore interferon α / β binding proteins were targeted

that are capable of modulating the activity of interferon-.

alpha. subtypes as well as interferon-β.

REFERENCE COUNT: 18 THERE ARE 18 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L4 ANSWER 19 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1996:91925 CAPLUS

DOCUMENT NUMBER: 124:139220

TITLE: Interferon-alpha/beta binding protein, its preparation and use

INVENTOR(S): Cohen, Batya; Novick, Daniela; Rubinstein, Menachem

PATENT ASSIGNEE(S): Israel

SOURCE: Can. Pat. Appl., 85 pp.

CODEN: CPXXEB

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
CA 2141747	 Δ1	19950808	CA 1995-2141747	19950203
	A	19950817		
	B2	19980312	NO 1999 11410	19330127
FI 9500516	. A	19950808	FI 1995-516	19950206
NO 9500420	A	19950808		
	B1	20050523		17733100
	A2	19951011	EP 1995-101560	19950206
EP 676413	A3	19960403		
EP 676413	B1	20050105	·	
R: AT, BE, CH,	DE, DK	, ES, FR,	GB, GR, IE, IT, LI, L	U, MC, NL, PT, SE
RU 2232811	C2 ·	20040720	RU 1995-101848	19950206
AT 286509	T	20050115	AT 1995-101560	· 19950206
PT 676413	T	20050531	PT 1995-101560	19950206
ES 2236696	T 3	20050716	ES 1995-101560	19950206
CN 1109505	Α	19951004	CN 1995-100194	19950207
ZA 9500968	Α	19951010	ZA 1995-968	19950207
JP 07298886	Α	19951114	JP 1995-43539	19950207
	B2	20050713		
JP 2005200422	Α	20050728		
PRIORITY APPLN. INFO.:		,	IL 1994-108584	
		•	JP 1995-43539	A3 19950207

AB Type I interferons (IFN- α and IFN- β and IFN- ω) are a family of cytokines usually defined by their ability to confer resistance to viral infections. There are pathol. situations, related to these cytokines where neutralization of interferon activity may be beneficial to the patient. Cytokine binding proteins (soluble cytokine receptors) correspond to the extracellular ligand binding domains of their resp. cell surface cytokine receptors. They are derived either by alternative splicing of pre-mRNA common to the cell surface receptor, or by proteolytic cleavage of the cell surface receptor. Therefore interferon α / β binding proteins were targeted that are capable of modulating the activity of interferon-. alpha. subtypes as well as interferon- β . Cloning of DNA mols. encoding these proteins and expression in host cells and antibodies against these proteins is discussed.

L4 ANSWER 20 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1995:332388 CAPLUS

DOCUMENT NUMBER: 122:104011

TITLE: Resistance of recombinant proteins to

proteolysis during folding and in the folded

state

AUTHOR(S): Fountoulakis, Michael

CORPORATE SOURCE: Dep. of Biology, F. Hoffmann-La Roche Ltd., Basel,

CH-4002, Switz.

SOURCE: Journal of Chemical Technology & Biotechnology (1995),

62(1), 81-90

CODEN: JCTBED; ISSN: 0268-2575

PUBLISHER: Wiley
DOCUMENT TYPE: Journal
LANGUAGE: English

Protein purification often involves the use of denaturing agents for solubilization. During refolding, following removal of the denaturants, the proteins of interest are exposed to proteases present in the expression system. Here the resistance of selected recombinant proteins to three widely used proteolytic enzymes, trypsin (EC 3.4.21.4), proteinase K (EC 3.4.21.14) and endoproteinase Glu-C (EC 3.4.21.19), was investigated during folding and in the folded state. Target proteins and protease mixts. were denatured in 8 mol dm-3 urea and the proteins were allowed to refold by removal of the urea by dialysis. The proteolytic products were analyzed by sodium dodecyl sulfate-polyacrylamide gels and protein digestion during folding was compared with the digestion under similar conditions in physiol. buffer. Depending on the folding state of the proteins, the proteases had different effects on the substrates. During folding, certain recombinant proteins were more efficiently digested by trypsin and, in particular, by proteinase K in comparison with digestion in the folded state, while other protein substrates were more resistant to proteolytic degradation in a denatured or partially denatured state than their folded counterparts. Incubation of most substrate proteins with endoproteinase Glu-C yielded kinetics of digestion that were essentially similar for both partially folded and unfolded substrates. The results reported may be useful for protection of sensitive proteins and in studies of protein folding mechanisms.

L4 ANSWER 21 OF 21 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1988:144742 CAPLUS

DOCUMENT NUMBER: 108:144742

TITLE: Identification and partial characterization of a novel

protease in Saccharomyces cerevisiae which cleaves the

peptide bond between residues 22 and 23 in $\alpha\text{-interferon},$ and identification of an

α-interferon resistant to said

proteolysis

INVENTOR(S): O'Loughlin, John T.

PATENT ASSIGNEE(S): Interferon Sciences, Inc., USA

SOURCE: Eur. Pat. Appl., 20 pp.

CODEN: EPXXDW

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 240224	A2	19871007	EP 1987-302519	19870324
EP 240224	A3	19890201		
R: AT, BE, CH,	DE, ES,	FR, GB, GR	, IT, LI, LU, NL, SE	
DK 8701614	Α	19871001	DK 1987-1614	19870330
CN 87102497	A	19871111	CN 1987-102497	19870330
JP 62296892	Α	19871224	JP 1987-74566	19870330

PRIORITY APPLN. INFO.:

US 1986-845937

A 19860331

AB A novel S. cerevisiae protease cleaves α -interferons between basic amino acids at positions 22 and 23, but cleavage does not occur if residue 22 is serine. A recombinant interferon α with serine, threonine, asparagine, glutamine, or glycine at position 22 could be produced intact in a microorganism whose primary proteolytic activity against the natural species is at that site. The protease was partially purified. from a protease-deficient PEP 3-4 S. cerevisiae mutant. It was membrane-bound and activated by the Triton X-100 present during cell lysis. Recombinant interferons α -1, α -2, and α -8 were all incubated with the protease. Both α -2 and α -8 were cleaved between amino acids 22 and 23 (which were Arg-Lys and Arg-Arg, resp.), but α -1 (Ser-Arg) was not.

=> Log off H
SESSION WILL BE HELD FOR 120 MINUTES
STN INTERNATIONAL SESSION SUSPENDED AT 13:58:52 ON 25 MAR 2007

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID: SSPTAEGS1646

PASSWORD:

* * * * * * RECONNECTED TO STN INTERNATIONAL * * * * * * SESSION RESUMED IN FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE' AT 14:18:26 ON 25 MAR 2007
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FILE 'CAPLUS' ENTERED AT 14:18:26 ON 25 MAR 2007
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FILE 'MEDLINE' ENTERED AT 14:18:26 ON 25 MAR 2007

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(FILE 'HOME' ENTERED AT 13:34:30 ON 25 MAR 2007)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE' ENTERED AT 13:34:48 ON 25 MAR 2007
L1 70256 S (INTERFERON ALPHA) OR IFN-ALPHA AND (MUTEIN OR VARIANT OR MUT
L2 248 S L1 AND PROTEOL?
L3 142 DUP REM L2 (106 DUPLICATES REMOVED)
L4 21 S L3 AND RESISTANCE

=> S L1 AND (IFN -alpha 2b)

L5 2141 L1 AND (IFN -ALPHA 2B)

=> S L5 AND Proteol?

L6 7 L5 AND PROTEOL?

L7 3 DUP REM L6 (4 DUPLICATES REMOVED)

=> D ti L7 1-3

L7 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2007 ACS on STN

TI High throughput directed evolution of proteins and peptides using two-dimensional rational mutagenesis scanning

L7 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 1

TI Identification of a linear epitope of interferon-.alpha .2b recognized by neutralizing monoclonal antibodies

L7 ANSWER 3 OF 3 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 2

TI NATURAL HUMAN INTERFERON-ALPHA-2 IS O-GLYCOSYLATED.

=> D Ibib ABs L7 2,3

L7 ANSWER 2 OF 3 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 1

ACCESSION NUMBER: 1999:656689 CAPLUS

DOCUMENT NUMBER: 132:11491

TITLE: Identification of a linear epitope of

interferon-α 2b recognized by

neutralizing monoclonal antibodies

AUTHOR(S): Blank, Viviana C.; Sterin-Prync, Aida; Retegui, Lilia;

Vidal, Alejandro; Criscuolo, Marcelo; Roguin, Leonor

Ρ.

CORPORATE SOURCE: Instituto de Quimica y Fisicoquimica Biologicas

(UBA-CONICET), Facultad de Farmacia y Bioquimica,

Buenos Aires, 1113, Argent.

SOURCE: European Journal of Biochemistry (1999), 265(1), 11-19

CODEN: EJBCAI; ISSN: 0014-2956

PUBLISHER: Blackwell Science Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

AB Four monoclonal antibodies (mAbs) directed against the recombinant human interferon- α 2b (IFN- α

2b) were used as probes to study the interaction of the IFN mol. to its receptors. The [125I]IFN-.alpha.2b

binding to immobilized mAbs was completely inhibited by IFN-.

alpha.2b and IFN- α 2a but neither IFN β nor

IFNγ showed any effect. Gel-filtration HPLC of the immune complexes

formed by incubating [125I] IFN-.alpha.2b

with paired mAbs revealed the lack of simultaneous binding of two different antibodies to the tracer, suggesting that all mAbs recognize the same IFN antigenic domain. Furthermore, the mAbs were also able to

neutralize the IFN-.alpha.2b anti-viral and

anti-proliferative activities as well as [1251] IFN-.

alpha.2b binding to WISH cell-membranes. As [1251] mAbs did not recognize IFN exposed epitopes in the IFN:receptor complexes, mAb induction of a conformational change in the IFN binding domain impairing its binding to receptors was considered unlikely. To identify the IFN

region recognized by mAbs, IFN-.alpha.2b was

digested with different proteolytic enzymes. Immunoreactivity of the resulting peptides was examined by Western blot and their sequences were established by Edman degradation after blotting to poly(vinylidene difluoride) membranes. Data obtained indicated that the smallest immunoreactive region recognized by mAbs consisted of residues 107-132 or 107-146. As this zone includes the sequence 123-140, which has been involved in the binding to receptors, and the authors' mAbs did not show an allosteric behavior, it is concluded that they are directed to

overlapping epitopes located close to or even included in the IFN binding domain.

REFERENCE COUNT: 47 THERE ARE 47 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L7 ANSWER 3 OF 3 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN

DUPLICATE 2

ACCESSION NUMBER: 1991:360709 BIOSIS

DOCUMENT NUMBER: PREV199192048934; BA92:48934

TITLE: NATURAL HUMAN INTERFERON-ALPHA-2 IS

O-GLYCOSYLATED.

AUTHOR (S): ADOLF G [Reprint author]; KALSNER I; AHORN H; MAURER-FOGY

I; CANTELL K

CORPORATE SOURCE: ERNST-BOEHRINGER-INST, ARZNEIMITTELFORSCHUNG, BENDER AND CO

BES MBH, DR BOEHRINGER-GASSE 5-11, A-1121 VIENNA, AUSTRIA

Biochemical Journal, (1991) Vol. 276, No. 2, pp. 511-518. SOURCE:

ISSN: 0264-6021.

DOCUMENT TYPE: Article FILE SEGMENT: BA LANGUAGE: ENGLISH

ENTRY DATE: Entered STN: 13 Aug 1991

Last Updated on STN: 13 Aug 1991

Natural human interferon α 2 (IFN- α 2) was

isolated from a preparation of partially purified human leucocyte IFN by monoclonal-antibody immunoaffinity chromatography. The purified protein had a specific activity of 1.5 + 108 i.u./mg; it was estimated to constitute 10-20% of the total antiviral activity of leucocyte IFN. N-Terminal amino-acid-sequence analysis identified the subspecies IFN-.alpha.2b and/or IFN- α 2c, whereas not detectable. The structure of natural IFN- $\alpha 2$ was found to differ from that of its recombinant (Escherichia coli-derived) equivalent. First, reverse-phase h.p.l.c. showed that natural IFN- α 2 was signficiantly more hydrophilic than expected. Secondly, the apparent molecular mass of the natural protein determined by SDS/PAGE was higher than that of recombinant IFN- $\alpha 2$; incubation under mild alkaline conditions known to eliminate O-linked carbohydrates resulted in a reduction of the apparent molecular mass to that of the recombinant protein. On sequence analysis of proteolytic peptides, Thr-106 was found to be modified. These results suggested that Thr-106 of natural IFN- α 2 carries O-linked carbohydrates. Reverse-phase h.p.l.c. as well as SDS/PAGE of natural IFN- α 2 showed that glycosylation is heterogeneous. For characterization of the carbohydrate moieties, the protein was treated with neuraminidase and/or O-glycanase and analysed by gel electrophoresis; in addition, glycopeptides obtained by proteinase digestion and separated by h.p.l.c. were characterized by sequence analysis and m.s. Further information on the composition of the glycans was obtained by monosaccharide analysis. The results indicate that natural IFN-α2 contains the disaccharide galactosyl-Nacetylgalactosamine (Gal-GalNAc) linked to the Thr-106. In part of the molecules, this core carbohydrate carries $(\alpha-)N$ -acetylneuraminic acid, whereas a disaccharide, probably N-acetyl-lactosamine, is bound to Gal-GalNAc in another proportion of the protein. Further glycosylation isomers are present in small amounts. As IFN- $\alpha 2$ is the only IFN- α species with a threonine residue at position 106, it may represent the only O-glycosylated human IFN- α protein.

=> D Hist

(FILE 'HOME' ENTERED AT 13:34:30 ON 25 MAR 2007)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE' ENTERED AT 13:34:48 ON 25 MAR 2007 L1 70256 S (INTERFERON ALPHA) OR IFN-ALPHA AND (MUTEIN OR VARIANT OR MUT

248 S L1 AND PROTEOL? L2

L3 142 DUP REM L2 (106 DUPLICATES REMOVED)

L4 21 S L3 AND RESISTANCE

L5 2141 S L1 AND (IFN -ALPHA 2B) L6 7 S L5 AND PROTEOL?

L7 3 DUP REM L6 (4 DUPLICATES REMOVED)

=> S L5 AND glycosyl?

L8 21 L5 AND GLYCOSYL?

=> Dup Rem. L8

PROCESSING COMPLETED FOR L8

L9 12 DUP REM L8 (9 DUPLICATES REMOVED)

- => D Ti L9 1-12
- L9 ANSWER 1 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 1
- TI Treatment of interferon- α for chronic hepatitis
- L9 ANSWER 2 OF 12 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 2
- TI GlycoPEGylation of recombinant therapeutic proteins produced in Escherichia coli.
- L9 ANSWER 3 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Study on mechanism of interferon treating pathological scars
- L9 ANSWER 4 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN
- TI High throughput directed evolution of proteins and peptides using two-dimensional rational mutagenesis scanning
- L9 ANSWER 5 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Methods and compositions relating to isoleucine boroproline compounds
- L9 ANSWER 6 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Providing natural allelic variants of interferon .alpha . as therapeutic agents with high therapeutic index
- L9 ANSWER 7 OF 12 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
- TI In vitro O-glycosylation of E. coli-produced therapeutic proteins using recombinant glycosyltransferases.
- L9 ANSWER 8 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 3
- TI Structural characterization of N-linked and O-linked oligosaccharides derived from interferon- α 2b and interferon- α 14c produced by Sendai-virus-induced human peripheral blood leukocytes
- L9 ANSWER 9 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 4
- TI Identification of nine interferon- α subtypes produced by Sendai virus-induced human peripheral blood leukocytes
- L9 ANSWER 10 OF 12 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
- TI Carbohydrate composition of natural source human-leukocyte derived interferon-alphan3.
- L9 ANSWER 11 OF 12 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 5
- TI Expression and purification of recombinant, glycosylated human interferon alpha 2b in murine myeloma NSo cells.
- L9 ANSWER 12 OF 12 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 6
- TI NATURAL HUMAN INTERFERON-ALPHA-2 IS O-GLYCOSYLATED.

L9 ANSWER 1 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 1

ACCESSION NUMBER: 2006:507346 CAPLUS

DOCUMENT NUMBER: 145:416104

TITLE: Treatment of interferon- α for chronic hepatitis C

AUTHOR(S): Moriyama, Mitsuhiko; Arakawa, Yasuyuki

CORPORATE SOURCE: Division of Gastroenterology and Hepatology,

Department of Medicine, Nihon University School of Medicine, Itabashi-ku, Tokyo, 173-8610, Japan

SOURCE: Expert Opinion on Pharmacotherapy (2006), 7(9),

1163-1179

CODEN: EOPHF7; ISSN: 1465-6566

PUBLISHER: Informa Healthcare
DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

AB A review. Combination therapy with polyethylene glycosylated

IFN-α2a or IFN-.alpha.2b and ribavirin

is currently the standard therapy for chronic hepatitis C. However, even with this therapy, hepatitis C virus cannot be eradicated in 50% of patients with refractory chronic hepatitis C. In addition, withdrawal or dose reduction occurs in .apprx. 40% of patients due to adverse effects. This treatment is also a contraindication in some patients, such as in patients with coexisting diseases or in elderly patients. For these patients, standard IFN- α monotherapy is even safer and more effective. In patients with chronic hepatitis C, IFN- α monotherapy results in a significant increase in the cumulative survival rate by suppressing the progression to hepatocellular carcinoma or liver failure. In addition, other efficacious therapeutic regimens have been employed, such as prolonged administration of standard IFN- α in elderly patients; prolonged low-dose continuous administration in patients with decompensated cirrhosis or hepatocellular carcinoma postoperative patients; and combination therapy with 5-fluorouracil and standard IFN- α for advanced hepatocellular carcinoma.

5-fluorouracil and standard IFN- α for advanced hepatocellular carcinoma. Monotherapy with standard IFN- α should thus be recognized as one of the

important therapeutic strategies for chronic hepatitis C.

REFERENCE COUNT: 84 THERE ARE 84 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 2 OF 12 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN

DUPLICATE 2

ACCESSION NUMBER: 2006:544386 BIOSIS DOCUMENT NUMBER: PREV200600541473

TITLE: GlycoPEGylation of recombinant therapeutic proteins

produced in Escherichia coli.

AUTHOR(S): DeFrees, Shawn; Wang, Zhi-Guang; Xing, Ruye; Scott, Arthur

E.; Wang, Jin; Zopf, David [Reprint Author]; Gouty,

Dominique L.; Sjoberg, Eric R.; Panneerselvam, Krishnasamy; Brinkman-Van der Linden, Els C. M.; Bayer, Robert J.; Tarp,

Mads A.; Clausen, Henrik

CORPORATE SOURCE: Neose Technol Inc, 102 Witmer Rd Dr, Horsham, PA 19044 USA

dzopf@neose.com

SOURCE: Glycobiology, (SEP 2006) Vol. 16, No. 9, pp. 833-843.

ISSN: 0959-6658.

DOCUMENT TYPE:

Article English

LANGUAGE: Engli

ENTRY DATE: Entered STN: 18 Oct 2006

Last Updated on STN: 18 Oct 2006

AB Covalent attachment of polyethylene glycol, PEGylation, has been shown to prolong the half-life and enhance the pharmacodynamics of therapeutic proteins. Current methods for PEGylation, which rely on chemical conjugation through reactive groups on amino acids, often generate isoforms in which PEG is attached at sites that interfere with bioactivity. Here, we present a novel strategy for site-directed PEGylation using glycosyltransferases to attach PEG to

O-glycans. The process involves enzymatic GaINAc glycosylation at specific serine and threonine residues in proteins expressed without glycosylation in Escherichia coli, followed by enzymatic transfer of sialic acid conjugated with PEG to the introduced GatNAc residues. The strategy was applied to three therapeutic polypeptides, granulocyte colony stimulating factor (G-CSF), interferon-alpha2b (IFNalpha 2b), and granulocyte/macrophage colony stimulating factor (GM-CSF), which are currently in clinical use.

ANSWER 3 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2006:714620 CAPLUS

DOCUMENT NUMBER: 146:272215

TITLE: Study on mechanism of interferon treating pathological

scars

Lu, Xin-an; Xu, Ming; Shen, Guo-liang; Lin, Wei; Zhao, AUTHOR (S):

Xiao-vu

Dept of Burn and Plastic Surgery, The First Hospital CORPORATE SOURCE:

Affiliated to Suzhou University, Jiangsu Suzhou,

215006, Peop. Rep. China

SOURCE: Suzhou Daxue Xuebao, Yixueban (2005), 25(6),

1091-1093, 1103

CODEN: SDXYC2; ISSN: 1673-0399

Suzhou Daxue Chubanshe PUBLISHER:

DOCUMENT TYPE: Journal LANGUAGE: Chinese

This paper studied the effect of interferon (IFN) on transforming growth

factor- β 1 (TGF- β 1), matrix metalloprotease-1 (MMP-1),

platelet-derived growth factor-BB (PDGF-BB), and glycosyltransferase (ppGalNAc-T2) in fibroblasts of pathol. scars and the mechanism of interferon on pathol. scars. The fibroblasts of pathol. scars were cultured by the method of tissue culture and were

randomized in 3 groups: control (0.9% sodium chloride), low concentration (100

u/mL IFN α -2b), and high concentration (10000

u/mL IFN α -2b). The expression of

 $TGF-\beta1$, MMP-1, PDGF-BB, and ppGalNAc-T2 were analyzed by RT-PCR in each group. The results showed that after treating cultured fibroblasts of pathol. scars with 100 u/mL IFN α -2b

and 10000 u/mL IFN α -2b, the expression of TGF- β 1, PDGF-BB, and ppGalNAc-T2 mRNA were lower than that in the control group, and the expression of MMP-1 mRNA was higher than that of control group. The result was significantly different and was related with the concentration in the IFN α -

2b. In conclusion, the cause of good effect of IFN . alpha.-2b on inhibiting fibroblast of pathol. scars may

relate with some kinds of cellular factors, such as TGF-β1 and PDGF-BB.

ANSWER 5 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 2004:41226 CAPLUS

DOCUMENT NUMBER: 140:105321

TITLE: Methods and compositions relating to isoleucine

boroproline compounds

INVENTOR(S): Adams, Sharlene; Miller, Glenn T.; Jesson, Michael I.; .

Jones, Barry

PATENT ASSIGNEE(S): Point Therapeutics, Inc., USA

PCT Int. Appl., 152 pp. SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE ----------------WO 2004004658 A2 20040115 WO 2003-US21405 20030709

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WO 2004004658
                           A3
                                 20050804
             AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM,
             PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN,
             TR, TT, TZ, UA, UG, UZ, VC, VN, YU, ZA, ZW
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES,
             FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                           A1
                                  20040115
                                            CA 2003-2491466
                                                                       20030709
     CA 2491466
                                 20040123
                                                                       20030709
     AU 2003265264
                           A1
                                              AU 2003-265264
     US 2004077601
                           A1
                                 20040422
                                              US 2003-616694
                                                                       20030709
                           A1
                                 20050421
                                              US 2003-616409
                                                                       20030709
     US 2005084490
                           A2
                                  20050928
                                             EP 2003-763380
                                                                       20030709
     EP 1578434
            AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
                          Т
                                 20060302
                                             JP 2004-562634
                                                                       20030709
     JP 2006507352
                           Α
                                 20060712
                                              CN 2003-821282
                                                                       20030709
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                           Α
                                 20060830 · CN 2003-821281
                                                                       20030709
     CN 1826129
     IN 2005KN00151
                           Α
                                  20050916
                                              IN 2005-KN151
                                                                       20050208
                                              US 2002-394856P
                                                                    P 20020709
PRIORITY APPLN. INFO.:
                                              US 2002-414978P
                                                                    P 20021001
                                              US 2003-466435P
                                                                    P
                                                                       20030428
                                              WO 2003-US21405
                                                                    W 20030709
OTHER SOURCE(S):
                          MARPAT 140:105321
     A method for treating subjects with, inter alia, abnormal cell
     proliferation or infectious disease using agents of formula (I,
     AmNHCH(CH(CH3)CH2CH3)COA1R) (where Am and Al are amino acids and R =
     organo boronates, organo phosphonates, fluoroalkyl ketones, alphaketos,
     N-peptiolyl-O-(acylhydroxylamines), azapeptides, azetidines, fluoroolefins
     dipeptide isosteres, peptidyl (\alpha-aminoalkyl) phosphonate esters,
     aminoacyl pyrrolidine-2-nitriles and 4-cyanothiazolidides) is claimed.
     Methods for stimulating an immune response using the compds. of the
     invention are also claimed. Compns. containing Ile-boroPro compds. are also
     provided as are kits containing the compns. The invention embraces the use of
     these compds. alone or in combination with other therapeutic agents.
     ANSWER 6 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN
L9
                          2004:39591 CAPLUS
ACCESSION NUMBER:
                          140:92604
DOCUMENT NUMBER:
                          Providing natural allelic variants of
TITLE:
                          interferon \alpha as therapeutic
                          agents with high therapeutic index
                          Escary, Jean-Louis
INVENTOR(S):
PATENT ASSIGNEE(S):
                          Fr.
                          U.S. Pat. Appl. Publ., 16 pp.
SOURCE:
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KIND
PATENT NO.
                          DATE
                                      APPLICATION NO.
                                                            DATE
                                      -----
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                          _ - - - - - -
                          20040115
                                     US 2002-315493
                                                            20021210
US 2004009161
                    A1
                          20040512
                                     EP 2002-292787
EP 1418428
                    A1
                                                            20021107
    R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
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CA 2413981
                    A1
CA 2504980
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                                      CA 2003-2504980
                                                            20031106
                    Α1
WO 2004042394
                    A2
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                                      WO 2003-EP13695
                                                            20031106
WO 2004042394
                    A3
                          20040715
    W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
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CODEN: USXXCO

Patent

1

English

DOCUMENT TYPE:

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

LANGUAGE:

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CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE,
              GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ,
              OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM,
              TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM,
                                                                       zw
         RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK,
              TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
     AU 2003294782
                            A1
                                   20040607
                                                AU 2003-294782
                                                                          20031106
                                   20050810
                                                EP 2003-785733
     EP 1561105
                            A2
                                                                        20031106
              AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
              IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
     US 2006094641
                            A1
                                   20060504
                                                US 2005-534098
                                                                          20050506
PRIORITY APPLN. INFO .:
                                                EP 2002-292787
                                                                      Α
                                                                         20021107
                                                US 2002-315493
                                                                      Α
                                                                          20021210
                                                WO 2003-EP13695
                                                                      W
                                                                          20031106
     Disclosed are methods for identifying and providing new therapeutic
AB
     agent(s) by selecting at least one polypeptide encoded by a natural
     allelic variant of one preselected gene having a therapeutic potential and
     determining the therapeutic index of the selected polypeptide(s) and retaining
     as therapeutic agent(s) those polypeptide(s) whose therapeutic index is
     higher than that of a reference agent. The invention is illustrated by tests
     performed on the polypeptides encoded by natural allelic variants of 3
     genes belonging to the interferon \alpha gene family
     and representing: C122S IFN\alpha-5; G45R IFN\alpha-17; and Q114H/V127D
     IFN\alpha-21 and K179E IFN\alpha-21. The polypeptides encoded by the natural allelic variants of IFN\alpha are subjected to several activity
     tests to determine their therapeutic suitability and are also compared with the
     product on the market, IFN\alpha -2\bar{b} (Intron
          The antiproliferative activities of the above variants were performed
     in tests on human lymphoblasts (Daudi cells) and their antiviral
     activities were evaluated in both virus-infected cell cultures (human WISH
     cells infection with vascular stomatitis virus) and mouse models
     (encephalomyocarditis virus and Friend erythroleukemia virus) of viral
     infection. The immunomodulatory activities of the above variants were
     tested on human dendritic cell maturation and a safety pharmacol. study
     was performed in conscious Rhesus monkeys.
     ANSWER 7 OF 12 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
T.9
ACCESSION NUMBER:
                      2005:321840 BIOSIS
DOCUMENT NUMBER:
                      PREV200510111622
TITLE:
                      In vitro O-glycosylation of E. coli-produced
                      therapeutic proteins using recombinant
                      glycosyltransferases.
                      Defrees, Shawn [Reprint Author]; Wang, Zhi-Guang; Scott,
AUTHOR (S):
                      Arthur E.; Wang, Jin; Xing, Ruye; Zopf, David; Gouty,
                      Dominique L.; Sjoberg, Eric R.; Panneerselvam, Krishnasamy;
                      Brinkman-Van der Linden, Els C. M.; Bayer, Robert J.; Tarp,
                      Mads A.; Clausen, Henrik
CORPORATE SOURCE:
                      Neose Technol Inc, Horsham, PA USA
                      Glycobiology, (NOV 2004) Vol. 14, No. 11, pp. 1086.
SOURCE:
                      Meeting Info.: Joint Meeting of the Society-for-
                      Glycobiology/Japanese-Society-for-Carbohydrate-Research.
                      Honolulu, HI, USA. November 17 -20, 2004. Soc Gylcobiol;
                      Japanese Soc Carbohydrate Res.
                      ISSN: 0959-6658.
DOCUMENT TYPE:
                      Conference; (Meeting)
                      Conference; Abstract; (Meeting Abstract)
LANGUAGE:
                      English
ENTRY DATE:
                      Entered STN: 25 Aug 2005
                      Last Updated on STN: 25 Aug 2005
```

ANSWER 8 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 3

1998:266317 CAPLUS

1,9

ACCESSION NUMBER:

DOCUMENT NUMBER: 129:39864

TITLE: Structural characterization of N-linked and O-linked

oligosaccharides derived from interferon-.

alpha.2b and interferon-.

alpha.14c produced by Sendai-virus-induced

human peripheral blood leukocytes

AUTHOR (S): Nyman, Tuula A.; Kalkkinen, Nisse; Tolo, Hannele;

Helin, Jari

CORPORATE SOURCE: Institute of Biotechnology, Protein Chemistry Lab.,

University of Helsinki, Finland

SOURCE: European Journal of Biochemistry (1998), 253(2),

485-493

CODEN: EJBCAI; ISSN: 0014-2956

PUBLISHER: Springer-Verlag

DOCUMENT TYPE: Journal LANGUAGE: English

The authors have previously isolated and partially characterized the components of a highly purified interferon- α

(IFN-α) preparation produced by Sendai-virus-induced human peripheral blood leukocytes. Nine IFN- α species were identified, and two of

these were glycosylated. Here, the authors isolated the

N-linked oligosaccharides of IFN- α 14c and the O-linked chains of

IFN-.alpha.2b, and the glycans were

characterized by electrospray tandem mass spectrometry and by specific glycosidase digestions monitored by matrix-assisted laser desorption ionization time of flight mass spectrometry. The IFN- α 14c N-glycans were shown to exhibit core-fucosylated biantennary glycans, with about 10% carrying an addnl. α 1,3-linked fucose unit at the antennae. The

IFN-.alpha.2b was shown to carry about 50%

core type-1 disialytetrasaccharides, 30% core type-1

monosialyltrisaccharides and 20% core type-2 monosialylpentasaccharides.

THERE ARE 45 CITED REFERENCES AVAILABLE FOR THIS REFERENCE COUNT: 45 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 9 OF 12 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 4

ACCESSION NUMBER: 1998:70613 CAPLUS

DOCUMENT NUMBER: 128:179185

TITLE: Identification of nine interferon-.

alpha. subtypes produced by Sendai

virus-induced human peripheral blood leukocytes Nyman, Tuula A.; Tolo, Hannele; Parkkinen, Jaakko;

Kalkkinen, Nisse

CORPORATE SOURCE: Institute of Biotechnology, Protein Chemistry

Laboratory, University of Helsinki, FIN-00014, Finland Biochemical Journal (1998), 329(2), 295-302

SOURCE:

CODEN: BIJOAK; ISSN: 0264-6021

PUBLISHER: Portland Press Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

AUTHOR(S):

The human interferon- α (IFN-.

alpha.) family is encoded by 13 different functional genes, and including all cloned sequence variants there are 28 potential

IFN- α proteins. To find out which of the

described sequences are expressed in normal human leukocytes, we have isolated and partly characterized the components of a highly purified

IFN- α preparation produced by Sendai virus-induced

human peripheral blood leukocytes. The identification protocol consisted of N-terminal sequencing and mass mapping of the proteins separated by reverse-phase HPLC and/or SDS/-PAGE. The highly purified leukocyte IFN- α preparation was found to contain at least nine

different IFN- α species: IFN-.

alpha.1a, IFN-.alpha.2b, IFN

 $-\alpha$ 4b, IFN- α 7a, IFN-.

alpha.8b, IFN- α 10a, IFN-.

alpha.14c, IFN- α 17b, and IFN-.

alpha.21b. IFN- α 1a was the major

subtype, comprising approx. 30% of total leukocyte IFN-.

alpha.. IFN- α 14c, the only subtype

containing potential N-glycosylation sites, was shown to be

glycosylated at Asn-72. Mol. mass determination of the intact proteins by

electrospray ionization MS showed that there are no other

post-translational modifications in the IFN- α subtypes than the glycosylation of IFN-.alpha

.2b and IFN- α 14c. Only one sequence

variant was found for each subtype, suggesting that the other described gene sequences represent allelic variants or mutations

that are more rarely found in the general population.

REFERENCE COUNT: 37 THERE ARE 37 CITED REFERENCES AVAILABLE FOR THIS

RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L9 ANSWER 10 OF 12 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on

STN

ACCESSION NUMBER: 1998:58374 BIOSIS DOCUMENT NUMBER: PREV199800058374

TITLE: Carbohydrate composition of natural source human-leukocyte

derived interferon-alphan3.

AUTHOR(S): Lawrynowicz, Witold J.; Lin, Xi; Lee, Shu-Ying;

Ferencz-Biro, Katalin; Liao, Mei-June

CORPORATE SOURCE: Interferon Sci. Inc., New Brunswick, NJ 08901, USA

SOURCE: Journal of Interferon and Cytokine Research, (Oct., 1997)

Vol. 17, No. SUPPL. 2, pp. S106. print.

Meeting Info.: Annual Meeting of the International Society

for Interferon and Cytokine Research. San Diego,

California, USA. October 19-24, 1997. International Society

for Interferon and Cytokine Research.

ISSN: 1079-9907.

DOCUMENT TYPE: Conference; (Meeting)

Conference; Abstract; (Meeting Abstract)

Conference; (Meeting Poster)

LANGUAGE:

English

ENTRY DATE: Entered STN: 30 Jan 1998

Last Updated on STN: 30 Jan 1998

L9 ANSWER 11 OF 12 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on

STN DUPLICATE 5

ACCESSION NUMBER: 1996:331039 BIOSIS DOCUMENT NUMBER: PREV199699053395

TITLE: Expression and purification of recombinant,

glycosylated human interferon

alpha 2b in murine myeloma NSo cells.

AUTHOR(S): Rossmann, Cornelia; Sharp, Nigel; Allen, Geoffrey; Gewert,

Dirk [Reprint author]

CORPORATE SOURCE: Cell Mol. Biol., Astra Draco AB, P.O. Box 34, S221 00 Lund,

Sweden

SOURCE: Protein Expression and Purification, (1996) Vol.

7, No. 4, pp. 335-342.

CODEN: PEXPEJ. ISSN: 1046-5928.

DOCUMENT TYPE:

Article

LANGUAGE:

English

ENTRY DATE:

Entered STN: 26 Jul 1996

Last Updated on STN: 27 Jul 1996

AB · We have expressed recombinant human interferon-alpha

-2b in mammalian cells and isolated cell lines constitutively secreting very high levels of biologically active protein. The expression system takes advantage of the strong human cytomegalovirus immediate early promoter in mouse myeloma NSo cells and glutamine synthetase as a selectable marker; spontaneous mutants with amplified gene copy numbers were selected by growth of primary transfectants in the presence of methionine sulfoximine. Using this procedure, we have isolated a recombinant NSo cell line which secretes human interferon at the rate of

20 mu-g/10-6 cells/24 h and accumulates up to 120 mu-g/ml (apprx 2.4 times 10-7 U/ml) following prolonged undiluted culture. The interferon (IFN) could be efficiently purified on a polyclonal bovine anti-human IFN-alpha specific antibody column and the glycosylation pattern was found to be similar to that of nonrecombinant IFN-alpha-2b purified from virus-induced human Namalwa cells. The biological activity of the recombinant material was indistinguishable from that of natural IFN from Namalwa cells, and the specific antiviral activity, as assayed on human HeLa cells challenged with encephalomyocarditis virus, was 2 times 10-8 IU/mg, similar to that of nonrecombinant IFN preparations. This represents the highest reported level of glycosylated, recombinant IFN expression in a stable mammalian system and is a significant advance in the large-scale production of these clinically important cytokines.

=> Log Off H
SESSION WILL BE HELD FOR 120 MINUTES
STN INTERNATIONAL SESSION SUSPENDED AT 14:29:15 ON 25 MAR 2007

Connecting via Winsock to STN

Welcome to STN International! Enter x:x

LOGINID:SSPTAEGS1646

PASSWORD:

* * * * * RECONNECTED TO STN INTERNATIONAL * * * * * * SESSION RESUMED IN FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE' AT 14:41:14 ON 25 MAR 2007
FILE 'BIOSIS' ENTERED AT 14:41:14 ON 25 MAR 2007
Copyright (c) 2007 The Thomson Corporation
FILE 'CAPLUS' ENTERED AT 14:41:14 ON 25 MAR 2007
COPYRIGHT (C) 2007 AMERICAN CHEMICAL SOCIETY (ACS)

FILE 'EMBASE' ENTERED AT 14:41:14 ON 25 MAR 2007 Copyright (c) 2007 Elsevier B.V. All rights reserved. FILE 'MEDLINE' ENTERED AT 14:41:14 ON 25 MAR 2007

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 178.38 178.59 DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION CA SUBSCRIBER PRICE -20.28 -20.28

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(FILE 'HOME' ENTERED AT 13:34:30 ON 25 MAR 2007)

FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE' ENTERED AT 13:34:48 ON 25 MAR 2007 L170256 S (INTERFERON ALPHA) OR IFN-ALPHA AND (MUTEIN OR VARIANT OR MUT 248 S L1 AND PROTEOL? L2L3 142 DUP REM L2 (106 DUPLICATES REMOVED) L421 S L3 AND RESISTANCE L5 2141 S L1 AND (IFN -ALPHA 2B) L6 7 S L5 AND PROTEOL? L73 DUP REM L6 (4 DUPLICATES REMOVED) 21 S L5 AND GLYCOSYL? L8 12 DUP REM L8 (9 DUPLICATES REMOVED) L9

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=> S L1 AND ((Increased Activity)(S)antiviral)
             0 L1 AND ((INCREASED ACTIVITY)(S) ANTIVIRAL)
L10
=> S L1 AND ((Increased Activity)(S)anti-proliferative)
L11
             O L1 AND ((INCREASED ACTIVITY)(S) ANTI-PROLIFERATIVE)
=> S L1 AND (Activity) (S) antiviral)
UNMATCHED RIGHT PARENTHESIS 'ANTIVIRAL)'
The number of right parentheses in a query must be equal to the
number of left parentheses.
=> S L1 AND (Activity(S)antiviral)
1.12
          3560 L1 AND (ACTIVITY(S) ANTIVIRAL)
=> S L1 AND (Activity(S)anti-proliferative)
L13
            93 L1 AND (ACTIVITY(S) ANTI-PROLIFERATIVE)
=> Dup Rem L12
PROCESSING IS APPROXIMATELY 56% COMPLETE FOR L12
PROCESSING COMPLETED FOR L12
           2040 DUP REM L12 (1520 DUPLICATES REMOVED)
=> S L14 AND ((IFN-alpha 2b) OR IFNalpha-2b)
L15
            71 L14 AND ((IFN-ALPHA 2B) OR IFNALPHA-2B)
=> S L13 AND ((IFN-alpha 2b) OR IFNalpha-2b)
             3 L13 AND ((IFN-ALPHA 2B) OR IFNALPHA-2B)
L16
=> Dup Rem L16
PROCESSING COMPLETED FOR L16
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L17
=> D Ti L17 1-2
    ANSWER 1 OF 2 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 1
L17
ΤI
     Identification of a linear epitope of interferon-.alpha
     .2b recognized by neutralizing monoclonal antibodies
1.17
    ANSWER 2 OF 2 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
     Natural killer cell activity against cultured melanoma cells: A
TТ
     dye-reduction technique with studies on augmented activity by interferon
     subtypes.
=> D Ibib L17 2
L17 ANSWER 2 OF 2 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
ACCESSION NUMBER:
                    1993:30348 BIOSIS
DOCUMENT NUMBER:
                    PREV199395018548
TITLE:
                    Natural killer cell activity against cultured melanoma
                    cells: A dye-reduction technique with studies on augmented
                    activity by interferon subtypes.
AUTHOR (S):
                    Losinno, Carmela; Wines, Bruce D.; Mackay, Terrance G.
                    Johns And Ian R. [Reprint author]
                    Cent. Mol. Biol. Med., Monash Univ., Clayton, Victoria
CORPORATE SOURCE:
                    3168, Australia
SOURCE:
                    Natural Immunity, (1992) Vol. 11, No. 4, pp. 215-224.
                    ISSN: 1018-8916.
DOCUMENT TYPE:
                    Article
```

English

Entered STN: 23 Dec 1992

Last Updated on STN: 24 Dec 1992

LANGUAGE:

ENTRY DATE:

- => D Ti L18 1-15
- L18 ANSWER 1 OF 15 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN Induction of APOBEC3 family proteins, a defensive maneuver underlying interferon-induced anti-HIV-1 activity.
- L18 ANSWER 2 OF 15 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN TGF-beta 1 mRNA expression in liver biopsy specimens and TGF-beta 1 serum levels in patients with chronic hepatitis C before and after antiviral therapy.
- L18 ANSWER 3 OF 15 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN Prolonging the half-life of human interferon-alpha2 in circulation:

 Design, preparation, and analysis of (2-sulfo-9-fluorenylmethoxycarbonyl)7-interferon-alpha2.
- L18 ANSWER 4 OF 15 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN .
 TI Hybrid (BDBB) interferon-alpha: Preformulation studies.
- L18 ANSWER 5 OF 15 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN TI NATURAL HUMAN INTERFERON-ALPHA-2 IS O-GLYCOSYLATED.
- L18 ANSWER 6 OF 15 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN BIOLOGIC ACTIVITY IN A FRAGMENT OF RECOMBINANT HUMAN INTERFERON ALPHA.
- L18 ANSWER 7 OF 15 CAPLUS COPYRIGHT 2007 ACS on STN
 TI High throughput directed evolution of proteins and peptides using two-dimensional rational mutagenesis scanning
- L18 ANSWER 8 OF 15 CAPLUS COPYRIGHT 2007 ACS on STN
 TI Proteolytic degradation of the recombinant target protein,
 interferon-\tau during its fermentative production in the methylotrophic
 yeast, Pichia pastoris
- L18 ANSWER 9 OF 15 CAPLUS COPYRIGHT 2007 ACS on STN
 TI The Nonstructural NS5A Protein of Hepatitis C Virus: An Expanding,
 Multifunctional Role in Enhancing Hepatitis C Virus Pathogenesis
- L18 ANSWER 10 OF 15 CAPLUS COPYRIGHT 2007 ACS on STN
 TI Long-acting cytokine derivatives and their pharmaceutical compositions
- L18 ANSWER 11 OF 15 CAPLUS COPYRIGHT 2007 ACS on STN
 TI Identification of a linear epitope of interferon-.alpha
 .2b recognized by neutralizing monoclonal antibodies
- L18 ANSWER 12 OF 15 CAPLUS COPYRIGHT 2007 ACS on STN
 TI Structural organization of the interferon molecules as precursors of immuno- and neuroactive oligopeptides
- L18 ANSWER 13 OF 15 CAPLUS COPYRIGHT 2007 ACS on STN TI Modified (1-28) beta interferons
- L18 ANSWER 14 OF 15 CAPLUS COPYRIGHT 2007 ACS on STN TI A new mass-spectrometric C-terminal sequencing technique finds a similarity between γ -interferon and $\alpha 2$ -interferon and identifies a proteolytically clipped γ -interferon that retains full antiviral activity
- L18 ANSWER 15 OF 15 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Interferon-mediated inhibition of production of Gazdar murine sarcoma virus, a retrovirus lacking env proteins and containing an uncleaved gag

AUTHOR (S):

SOURCE:

=> D Ibib Abs L18 3, 4, 13, 14

L18 ANSWER 3 OF 15 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN

ACCESSION NUMBER: 2001:147461 BIOSIS DOCUMENT NUMBER: PREV200100147461

TITLE: Prolonging the half-life of human interferon-alpha2 in

circulation: Design, preparation, and analysis of

(2-sulfo-9-fluorenylmethoxycarbonyl) 7-interferon-alpha2. Shechter, Yoram [Reprint author]; Preciado-Patt, Liana;

Schreiber, Gideon; Fridkin, Mati

CORPORATE SOURCE: Department of Biological Chemistry, Weizmann Institute of

Science, Rehovot, 76100, Israel

yoram.shechter@weizmann.ac.il; mati.fridkin@weizmann.ac.il Proceedings of the National Academy of Sciences of the United States of America, (January 30, 2001) Vol. 98, No.

3, pp. 1212-1217. print.

CODEN: PNASA6. ISSN: 0027-8424.

DOCUMENT TYPE: Article LANGUAGE: English

ENTRY DATE: Entered STN: 21 Mar 2001

Last Updated on STN: 15 Feb 2002

Polypeptide drugs are generally short-lived species in circulation. In this study, we have covalently linked seven moieties of 2-sulfo-9-fluorenylmethoxycarbonyl (FMS) to the amino groups of human interferon-alpha2. The derivative thus obtained (FMS7-IFN-alpha2) has apprxeq4% the biological potency and 33 +- 4% the receptor binding capacity of the native cytokine. Upon incubation, FMS7-IFN-alpha2 undergoes time-dependent spontaneous hydrolysis, generating active interferon with t1/2 values of 24 +- 2 h at pH 8.5 and 98 +- 10 h at pH When native IFN- alpha2 is intravenously administered to mice, circulating antiviral activity is maintained for a short duration and then declines with t1/2 = 4 +- 0.5 h, reaching undetectable values at apprxeq18 h after administration. With intravenously administered FMS7-IFN-alpha2, there is a lag period of 2 h, followed by a progressive elevation in circulating antiviral-active protein, which peaked at 20 h and declined with t1/2 = 35 + - 4 h. FMS7-IFN-alpha2 is resistant to alpha-chymotrypsin digest and to proteolytic inactivation by human serum proteases in vitro. We have thus introduced here an inactive IFN-alpha2 derivative, which is resistant to in situ inactivation and has the capability of slowly reverting to the native active protein at physiological conditions in vivo and in vitro. Having these attributes, FMS7-IFN-alpha2 maintains prolonged circulating antiviral activity in mice, exceeding 7-8 times the activity of intravenously administered native cytokine.

L18 ANSWER 4 OF 15 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN

ACCESSION NUMBER: 1999:521402 BIOSIS DOCUMENT NUMBER: PREV199900521402

TITLE: Hybrid (BDBB) interferon-alpha:

Preformulation studies.

AUTHOR(S): Allen, John D.; Bentley, David; Stringer, Rowan A.;

Lowther, Nicholas [Reprint author]

CORPORATE SOURCE: Drug Preformulation and Delivery Department, Ciba

Pharmaceuticals (now Novartis Horsham Research Centre), Wimblehurst Road, Horsham, West Sussex, RH12 5AB, UK International Journal of Pharmaceutics (Amsterdam), (Oct.

5, 1999) Vol. 187, No. 2, pp. 259-272. print.

CODEN: IJPHDE. ISSN: 0378-5173.

DOCUMENT TYPE: Article LANGUAGE: English

SOURCE:

ENTRY DATE: Entered STN: 3 Dec 1999

Last Updated on STN: 3 Dec 1999

AB A number of techniques, including RP-HPLC, HP-SEC and SDS-PAGE have been used in the delineation of degradative mechanisms of recombinant hybrid (BDBB) interferon-alpha (IFN-alpha) in the solution phase. Different degradation profiles are found according to medium pH. At pH 4.0 the major routes of degradation are via chemical transformation of the monomeric protein to a species which retains antiviral activity, and by self-proteolytic hydrolysis. At pH 7.6, methionine-oxidation is the major chemical degradative process. Protein aggregation is also a significant route of degradation at the higher pH. The results have assisted in a targeted preformulation screen of potentially stabilising excipients and possible parenteral solution dosage forms have been identified. Preliminary 'real-time' storage data confirm excellent chemical and physical stability of IFN-alpha in vehicles formulated at pH 7.6 or, especially, pH 4.0 under the proposed shelf conditions.

L18 ANSWER 13 OF 15 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1985:417791 CAPLUS

DOCUMENT NUMBER:

103:17791

TITLE:

Modified (1-28) beta interferons

INVENTOR(S):

Bell, Leslie D.; Boseley, Paul G.; Smith, John C.;

Houghton, Michael

PATENT ASSIGNEE(S):

G.D. Searle and Co., USA

Eur. Pat. Appl., 64 pp.

DOCUMENT TYPE:

CODEN: EPXXDW Patent

LANGUAGE:

SOURCE:

English

FAMILY ACC. NUM. COUNT: 4

PATENT INFORMATION:

PATENT NO.	KIND DAT	'E API	PLICATION NO.	DATE
EP 130566		50109 EP	1984-107458	19840628
		71028		
R: BE, CH, DE,			Ξ	
US 4738844			1984-623814	19840622
			1984-623601	19840622
US 4793995			1984-623815	19840622
US 4738845			1984-623894	19840625
AU 8429981			1984-29981	19840628
AU 577789		81006		
AU 8429982			1984-29982	19840628
AU 577790		81006		
AU 8429983			1984-29983 ·	19840628
AU 577791		81006		
AU 8429984			1984-29984	19840628
AU 577792		81006		
JP 60100599			1984-137079	19840702
JP 60105700			1984-137081	19840702
JP 60143000			1984-137080	19840702
JP 60214800			1984-137078	19840702
PRIORITY APPLN. INFO.:			1983-17880 A	
AR Pecombinant DNA mole	are cons			

Recombinant DNA mols. are constructed which encode modified human β -interferon (IFN- β) mols. The modification involves replacement by 3-28 amino acids of amino acids nos. 1-28, in some cases by amino acids 2-28 from α -interferon. Plasmid vectors for these modified IFN mols. are also prepared One modified IFN- β contains serine at position 16 in place of cysteine. Other IFNs contain α -IFN sequences. These modified interferons (designated group I IFNs) display some of the following properties; greater antiproliferative or antiviral activity, modified affinity for cell surface receptors, increased therapeutic index, increased stability in proteolysis, increased solubility in vivo, and greater ease of purification or recovery from bacterial exts. Pharmaceutical compns. containing these

modified mols. are used to treat viral infections, regulate cell growth (as an antineoplastic agent), or regulate the immune system. Thus, amino acids 1-28 were replaced in groups of 3-28 amino acids by the insertion of chemical synthesized oligodeoxyribonucleotide blocks. The oligodeoxyribonucleotides were prepared by the phosphoramidate method. Blocks (30-50 bases) were assembled by combining each phosphorylated component with equimolar amts. of the unphosphorylated oligomers from the complementary strand. Plasmid vectors were then used to clone the synthetic DNA fragments into the IFN- β -coding region. The vectors also contained the Escherichia coli trp promoter. The IFN- β formed by E. coli (IFNX414) had in vitro antiviral and antiproliferative activities .apprx.5-fold higher than those of IFN- β . Another recombinant IFN- β , IFNX401 had identical antiviral and immunostimulating activity to IFN- β but is 3 times more potent in its antiproliferative activity. Other group I IFNs prepared and characterized were IFNX412, 413, 421, and modified β -.

L18 ANSWER 14 OF 15 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1984:4461 CAPLUS

DOCUMENT NUMBER: 100:4461

TITLE:

A new mass-spectrometric C-terminal sequencing

technique finds a similarity between γ -interferon and α 2-interferon and identifies a proteolytically clipped γ -interferon that retains full antiviral

activity

AUTHOR(S): Rose, Keith; Simona, Marco G.; Offord, Robin E.;

Prior, Christopher P.; Otto, Berndt; Thatcher, David

R.

CORPORATE SOURCE: Dep. Biochim., Cent. Med. Univ., Geneva, 1211/4,

Switz.

SOURCE: Biochemical Journal (1983), 215(2), 273-7

CODEN: BIJOAK; ISSN: 0306-3275

DOCUMENT TYPE: LANGUAGE: Journal English

AB During peptide sequence mapping, it is difficult to obtain sequence information from the C-terminus; it is much easier to obtain sequence information from the N-terminus of a protein (Rose, K., et al, 1983). A novel mass-spectrometric technique is described here which permits identification of the C-terminal peptide of a protein. This technique involves the incorporation of 180 into all α-carboxy groups liberated during enzyme-catalyzed partial hydrolysis of the protein, followed by mass spectrometry to identify as the C-terminal peptide the only peptide that did not incorporate any 180. This technique was used to identify the true C-terminal tryptic peptide of a bacterially-produced (recombinant technol.) γ-interferon (human) and to distinguish it from a peptide produced by an anomalous tryptic cleavage. A closely similar sequence segment of bacterially produced α2-interferon undergoes an analogous cleavage. The C-terminus of a clipped γ-interferon that retains full antiviral activity also was identified by using the technique.

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          70256 S (INTERFERON ALPHA) OR IFN-ALPHA AND (MUTEIN OR VARIANT OR MUT
L1
L2
            248 S L1 AND PROTEOL?
            142 DUP REM L2 (106 DUPLICATES REMOVED)
L3
L4
             21 S L3 AND RESISTANCE
L5
           2141 S L1 AND (IFN -ALPHA 2B)
L6
              7 S L5 AND PROTEOL?
L7
             3 DUP REM L6 (4 DUPLICATES REMOVED)
L8
             21 S L5 AND GLYCOSYL?
L9
            12 DUP REM L8 (9 DUPLICATES REMOVED)
L10
             0 S L1 AND ((INCREASED ACTIVITY)(S)ANTIVIRAL)
L11
             0 S L1 AND ((INCREASED ACTIVITY)(S)ANTI-PROLIFERATIVE)
L12
           3560 S L1 AND (ACTIVITY(S)ANTIVIRAL)
            93 S L1 AND (ACTIVITY(S)ANTI-PROLIFERATIVE)
L13
           2040 DUP REM L12 (1520 DUPLICATES REMOVED)
L14
             71 S L14 AND ((IFN-ALPHA 2B) OR IFNALPHA-2B)
L15
L16
              3 S L13 AND ((IFN-ALPHA 2B) OR IFNALPHA-2B)
L17
              2 DUP REM L16 (1 DUPLICATE REMOVED)
             15 S L14 AND PROTEOL?
L18
=> S L1(P)Protease
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH
FIELD CODE - 'AND' OPERATOR ASSUMED 'L1(P) PROTEASE'
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH
FIELD CODE - 'AND' OPERATOR ASSUMED 'L2(P) PROTEASE'
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH
FIELD CODE - 'AND' OPERATOR ASSUMED 'L3(P) PROTEASE'
PROXIMITY OPERATOR LEVEL NOT CONSISTENT WITH
FIELD CODE - 'AND' OPERATOR ASSUMED 'L4(P) PROTEASE'
L19
           837 L1(P) PROTEASE
=> Dup Rem L19
PROCESSING COMPLETED FOR L19
L20
            588 DUP REM L19 (249 DUPLICATES REMOVED)
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=> D Ti

L20 ANSWER 1 OF 588 CAPLUS COPYRIGHT 2007 ACS on STN
TI Preparation of spiroisoxazoline-based peptidomimetics as inhibitors of

serine proteases, particularly HCV NS3-NS4A protease

- => S ((INTERFERON ALPHA) OR IFN-ALPHA)(P)protease L21 547 ((INTERFERON ALPHA) OR IFN-ALPHA)(P) PROTEASE
- => S L21 AND pd<=20020909 L22 290 L21 AND PD<=20020909
- => Dup Rem L22
 PROCESSING COMPLETED FOR L22
 L23 136 DUP REM L22 (154 DUPLICATES REMOVED)
- => D Ti 1-5
- L23 ANSWER 1 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
- TI AIDS-related Kaposi's sarcoma with chylothorax and pericardial involvement satisfactorily treated with liposomal doxorubicin.
- L23 ANSWER 2 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Preparation of recombinant protein as chaperon fusion protein
- L23 ANSWER 3 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Adhesion protein, protease, and protease inhibitor mutations and methods for diagnosis and treatment of epithelial cell adhesion-associated diseases
- L23 ANSWER 4 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Preparation of peptidomimetic protease inhibitors
- L23 ANSWER 5 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Preparation of novel peptides as NS3-serine protease inhibitors of hepatitis C virus
- => D Ti 1-136
- L23 ANSWER 1 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
- TI AIDS-related Kaposi's sarcoma with chylothorax and pericardial involvement satisfactorily treated with liposomal doxorubicin.
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- TI Preparation of recombinant protein as chaperon fusion protein
- L23 ANSWER 3 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Adhesion protein, protease, and protease inhibitor mutations and methods for diagnosis and treatment of epithelial cell adhesion-associated diseases
- L23 ANSWER 4 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Preparation of peptidomimetic protease inhibitors
- L23 ANSWER 5 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Preparation of novel peptides as NS3-serine protease inhibitors of hepatitis C virus
- L23 ANSWER 6 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Novel peptides as ns3-serine protease inhibitors of hepatitis C virus
- L23 ANSWER 7 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Preparation of novel imidazolidinones as NS3-serine protease inhibitors of hepatitis C virus

- L23 ANSWER 8 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Preparation of novel peptides as NS3-serine protease inhibitors of hepatitis C virus
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- TI Ambroxol suppresses influenza-virus proliferation in the mouse airway by increasing antiviral factor levels.
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 Original Title: Traitement de l'hepatite C.
- L23 ANSWER 14 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
- TI Adverse drug reaction update.
- L23 ANSWER 15 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- Proteolytic degradation of the recombinant target protein, interferon- τ during its fermentative production in the methylotrophic yeast, Pichia pastoris
- L23 ANSWER 16 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 3
- TI Management of protease inhibitor-associated hyperlipidemia
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- TI Monitoring of endogenous interferon-alpha and human herpesvirus 8 in HIV-infected patients with Kaposi's sarcoma.
- L23 ANSWER 18 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Preparation of macrocyclic NS3-serine protease inhibitors of hepatitis C virus comprising n-cyclic p2 moieties
- L23 ANSWER 19 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Preparation of peptides as inhibitors of serine proteases, particularly hepatitis C virus NS3 protease
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- L23 ANSWER 21 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 4
- TI Experimental and emerging therapies for chronic hepatitis C virus infection
- L23 ANSWER 22 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 5
- TI Prolonging the half-life of human interferon- $\alpha 2$ in circulation: design, preparation, and analysis of (2-sulfo-9-fluorenylmethoxycarbonyl)7-interferon- $\alpha 2$
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- TI Treatment with interferon-alpha (IFNalpha) of hepatitis C patients induces lower serum dipeptidyl peptidase IV activity, which is related to IFNalpha-induced depressive and anxiety symptoms and immune activation.
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- TI Therapeutic uses of protease inhibitors to modulate cellular pathways and immunity
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- TI RNase-L-dependent destabilization of interferon-induced mRNAs. A role for the 2-5A system in attenuation of the interferon response
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- TI Localization of a receptor nonapeptide with a possible role in the binding of the type I interferons.
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- TI [Coinfection with the hepatitis C virus and HIV: Current aspects].

 CO-INFECTION PAR LE VIRUS DE L'HEPATITE C ET LE VIRUS DE
 L'IMMUNODEFICIENCE HUMAINE: ASPECTS ACTUELS.
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- TI Suppression of hepatitis C virus in human immunodeficiency virus iron-loading anemia (HCV-HIV-ILA) patients with HAART and recombinant human erythropoietin (r-HuEPO).
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- TI STAT1 plays a protective role against the neurotoxic actions of chronic IFN-alpha production in the CNS.
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- TI Mastocytosis.

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- TI Obstruction of HIV-1 particle release by interferon-. alpha. occurs before viral protease processing and is independent of envelope glycoprotein
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- TI Obstruction of HIV-1 particle release by interferonalpha occurs before viral protease processing and is independent of envelope glycoprotein.
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- TI A phase II study of interferon-alpha, interleukin-2 and 5-fluorouracil in advanced renal carcinoma: Clinical data and laboratory evidence of protease activation.
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- TI Inhibition of human immunodeficiency virus type 1 replication in cytokine-stimulated monocytes/macrophages by combination therapy.
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- TI Structure and physicochemical properties of purified human leukocyte interferon (FPI-31)
- L23 ANSWER 91 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
- TI Interferon-alpha induces plasma interleukin-6 elevation in patients with chronic hepatitis C: Its abrogation by a serine protease inhibitor.
- L23 ANSWER 92 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 39
- TI Involvement of antipain-sensitive protease activity in suppression of UV-mutagenicity by human interferon-alpha

- L23 ANSWER 93 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 40
- TI A stress-regulated protein, GRP58, a member of thioredoxin superfamily, is a carnitine palmitoyltransferase isoenzyme.
- L23 ANSWER 94 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
- TI Inhibition of HIV-1 replication by convergent combination therapy in monocyte/macrophages.
- L23 ANSWER 95 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 41
- TI Regulation of neutrophil-derived IL-8: The role of prostaglandin E-2, dexamethasone, and IL-4.
- L23 ANSWER 96 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 42
- TI Inhibition of the protease of human immunodeficiency virus blocks replication and infectivity of the virus in chronically infected macrophages.
- L23 ANSWER 97 OF 136 EMBASE COPYRIGHT (c) 2007 Elsevier B.V. All rights reserved on STN
- TI New antiretroviral agents for the therapy of HIV type-1 infection.
- L23 ANSWER 98 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 43
- TI Damage of tracer erythropoietin results in erroneous estimation of concentration in mouse submaxillary gland
- L23 ANSWER 99 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 44
- TI Duplication of secretion signal sequences is deleterious for the secretion of human interferon $\alpha 4$ from Saccharomyces cerevisiae and Bacillus subtilis
- L23 ANSWER 100 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 45
- TI Inhibition of antigen-induced secretion in the rat jejunum by interferon alpha/beta.
- L23 ANSWER 101 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- TI In vitro processing of fusion proteins
- L23 ANSWER 102 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 46
- TI Human immunodeficiency virus type 1 (HIV-1) inhibitory interactions between protease inhibitor Ro 31-8959 and zidovudine, 2' 3'-dideoxycytidine, or recombinant interferon-alpha A against zidovudine-sensitive or -resistant HIV-1 in vitro.
- L23 ANSWER 103 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 47
- TI FAT-STORING CELLS OF THE RAT LIVER SYNTHESIZE AND SECRETE C1 ESTERASE INHIBITOR MODULATION BY CYTOKINES.
- L23 ANSWER 104 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 48
- TI DIFFERENTIAL INACTIVATION OF INTERFERONS BY A PROTEASE FROM HUMAN GRANULOCYTES.
- L23 ANSWER 105 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 49
- TI Rapid high level production and purification of recombinant murine and human interferons alpha from Escherichia coli.
- L23 ANSWER 106 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on

STN DUPLICATE 50

TI DIFFERENTIAL MODULATION OF TWO INTERFERON-ALPHA BINDING PROTEINS ON A HUMAN LYMPHOBLASTOID CELL LINE.

- L23 ANSWER 107 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 51
- TI INTERFERON INHIBITOR IN THE BLOOD OF PATIENTS WITH SYSTEMIC LUPUS ERYTHEMATOSUS.
- L23 ANSWER 108 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Glycosylated polypeptides for better thermostability and protease resistance
- L23 ANSWER 109 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 52
- TI STRUCTURAL DESIGN AND MOLECULAR EVOLUTION OF A CYTOKINE RECEPTOR SUPERFAMILY.
- L23 ANSWER 110 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 53
- TI INTERFERON GAMMA INCREASES IN-VITRO AND IN-VIVO EXPRESSION OF C1 INHIBITOR.
- L23 ANSWER 111 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 54
- TI A SENSITIVE TWO-SITE ENZYME IMMUNOASSAY FOR THE DETECTION OF RAT INTERFERON-GAMMA IN BIOLOGICAL FLUIDS.
- L23 ANSWER 112 OF 136 EMBASE COPYRIGHT (c) 2007 Elsevier B.V. All rights reserved on STN DUPLICATE 55
- TI Virologic and immunologic aspects of acquired immunodeficiency syndrome.
- L23 ANSWER 113 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Identification of actinophage VWB promoters and their use for expression of murine interferon alpha in Streptomyces venezuelae and S. lividans
- L23 ANSWER 114 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Inhibition of human natural killer cell activity by Legionella pneumophila protease
- L23 ANSWER 115 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 56
- TI SECRETORY EXPRESSION IN ESCHERICHIA-COLI AND BACILLUS-SUBTILIS OF HUMAN INTERFERON ALPHA GENES DIRECTED BY STAPHYLOKINASE SIGNALS.
- L23 ANSWER 116 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 57
- TI Mass spectrometric analysis of recombinant human α -2 interferon
- L23 ANSWER 117 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 58
- TI LOW TEMPERATURES STABILIZE INTERFERON ALPHA-2 AGAINST PROTEOLYSIS IN METHYLOPHILUS-METHYLOTROPHUS AND ESCHERICHIA-COLI.
- L23 ANSWER 118 OF 136 MEDLINE on STN
- TI [Monocyte-endothelium relations]. Relations monocytes-endothelium.
- L23 ANSWER 119 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 59
- TI CYTOSTATIC PRODUCTS RELEASED BY ACTIVATED MACROPHAGES UNRELATED TO INTERLEUKIN 1 TUMOR NECROSIS FACTOR ALPHA AND INTERFERON-ALPHA-BETA.
- L23 ANSWER 120 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 60
- TI SELECTIVE INDUCTION OF MONONUCLEAR PHAGOCYTES TO PRODUCE NEOPTERIN BY

INTERFERONS.

- ANSWER 121 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on DUPLICATE 61
- TI THE STABILITY OF NORMAL ABNORMAL AND GENETICALLY-ENGINEERED PROTEINS IN ESCHERICHIA-COLI STRAINS DEFICIENT IN THE LON-GENE PRODUCTS INTRACELLULAR PROTEASE LA.
- ANSWER 122 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN L23
- Plasmids which include promoters for bacteriocins adapted for expression of foreign polypeptides in Escherichia coli
- L23
- ANSWER 123 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN Identification and partial characterization of a novel protease in Saccharomyces cerevisiae which cleaves the peptide bond between residues 22 and 23 in α -interferon, and identification of an α-interferon resistant to said proteolysis
- ANSWER 124 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on L23 **DUPLICATE 62** STN
- INTERFERON-GAMMA IS A MAJOR REGULATOR OF C1-INHIBITOR SYNTHESIS BY HUMAN ΤI BLOOD MONOCYTES.
- L23 ANSWER 125 OF 136 MEDLINE on STN
- Production and function of the monocyte cytotoxic factor (MCF).
- L23 ANSWER 126 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on **DUPLICATE 63** STN
- HUMAN MONOCYTE OR RECOMBINANT INTERLEUKIN 1'S ARE SPECIFIC FOR THE TΙ SECRETION OF A METALLOPROTEINASE FROM CHONDROCYTES.
- L23 ANSWER 127 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on **DUPLICATE 64**
- INTRACELLULAR DEGRADATION OF RECOMBINANT PROTEINS IN RELATION TO THEIR ΤI LOCATION IN ESCHERICHIA-COLI CELLS.
- L23 ANSWER 128 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on **DUPLICATE 65**
- HUMAN INTERFERON GAMMA INCREASES ADHESION OF CULTURED CARCINOMA CELLS TO TI THE SUBSTRATUM.
- ANSWER 129 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on L23 DUPLICATE 66
- A LYMPHOKINE REGULATES EXPRESSION OF ALPHA-1 PROTEINASE INHIBITOR IN HUMAN TI MONOCYTES AND MACROPHAGES.
- ANSWER 130 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN L23
- TT Secretion of mature IFN- α 2 and accumulation of uncleaved precursor by Bacillus subtilis transformed with a hybrid α-amylase signal sequence-IFN-α2 gene
- ANSWER 131 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN L23
- An interferon-like activity released by human peripheral blood mononuclear cells treated with inhibitors of transcription: a new type of interferon?
- ANSWER 132 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN L23
- Interferon administration vehicles ΤI
- L23 ANSWER 133 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN **DUPLICATE 67**
- TI MONOCLONAL ANTIBODIES ELECTROPHORETICALLY TRANSFERRED FROM POLYACRYLAMIDE GELS RETAIN THEIR ABILITY TO BIND SPECIFIC ANTIGENS.
- L23 ANSWER 134 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 68

- TI ENZYME IMMUNOASSAY OF HUMAN INTERFERON-ALPHA USING MONOCLONAL ANTIBODY.
- L23 ANSWER 135 OF 136 MEDLINE on STN
- TI Interferon is a polypeptide hormone.
- L23 ANSWER 136 OF 136 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN DUPLICATE 69
- TI REDUCED SYNTHÉSIS OF PP-60 S-R-C AND EXPRESSION OF THE TRANSFORMATION RELATED PHENOTYPE IN INTERFERON TREATED ROUS SARCOMA VIRUS TRANSFORMED RAT CELLS.
- => S L23 AND (Interferon-alpha 2b) L24 5 L23 AND (INTERFERON-ALPHA 2B)
- => D Ti 1-5
- L24 ANSWER 1 OF 5 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN TI Treatment of hepatitis.
- L24 ANSWER 2 OF 5 BIOSIS COPYRIGHT (c) 2007 The Thomson Corporation on STN
- TI Treatment strategies for chronic hepatitis C: Update since the 1997
 National Institutes of Health Consensus Development Conference.
- L24 ANSWER 3 OF 5 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Preparation of recombinant protein as chaperon fusion protein
- L24 ANSWER 4 OF 5 CAPLUS COPYRIGHT 2007 ACS on STN
- TI Preparation of peptidomimetic protease inhibitors
- L24 ANSWER 5 OF 5 EMBASE COPYRIGHT (c) 2007 Elsevier B.V. All rights reserved on STN
- TI Mastocytosis.
- => Loggoff h
 LOGGOFF IS NOT A RECOGNIZED COMMAND
 The previous command name entered was not recognized by the system.
 For a list of commands available to you in the current file, enter
 "HELP COMMANDS" at an arrow prompt (=>).
- => log off h SESSION WILL BE HELD FOR 120 MINUTES STN INTERNATIONAL SESSION SUSPENDED AT 15:14:44 ON 25 MAR 2007

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COPYRIGHT (C) 2007 AMERICAN CHEMICAL SOCIETY (ACS)
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FULL ESTIMATED COST
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                                                                  340.80
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                                                                   TOTAL
                                                       ENTRY
                                                                 SESSION
CA SUBSCRIBER PRICE
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     (FILE 'HOME' ENTERED AT 13:34:30 ON 25 MAR 2007)
     FILE 'BIOSIS, CAPLUS, EMBASE, MEDLINE' ENTERED AT 13:34:48 ON 25 MAR 2007
          70256 S (INTERFERON ALPHA) OR IFN-ALPHA AND (MUTEIN OR VARIANT OR MUT
L1
            248 S L1 AND PROTEOL?
L2
L3
            142 DUP REM L2 (106 DUPLICATES REMOVED)
             21 S L3 AND RESISTANCE
L4
L5 .
           2141 S L1 AND (IFN -ALPHA 2B)
1.6
              7 S L5 AND PROTEOL?
1.7
              3 DUP REM L6 (4 DUPLICATES REMOVED)
T.R
             21 S L5 AND GLYCOSYL?
L9
             12 DUP REM L8 (9 DUPLICATES REMOVED)
L10
              0 S L1 AND ((INCREASED ACTIVITY)(S)ANTIVIRAL)
L11 '
              0 S L1 AND ((INCREASED ACTIVITY)(S)ANTI-PROLIFERATIVE)
L12
           3560 S L1 AND
                         (ACTIVITY(S)ANTIVIRAL)
L13
             93 S L1 AND (ACTIVITY(S)ANTI-PROLIFERATIVE)
L14
           2040 DUP REM L12 (1520 DUPLICATES REMOVED)
L15
             71 S L14 AND ((IFN-ALPHA 2B) OR IFNALPHA-2B)
              3 S L13 AND ((IFN-ALPHA 2B) OR IFNALPHA-2B)
L16
L17
              2 DUP REM L16 (1 DUPLICATE REMOVED)
L18
             15 S L14 AND PROTEOL?
            837 S L1(P)PROTEASE
L19
            588 DUP REM L19 (249 DUPLICATES REMOVED)
L20
            547 S ((INTERFERON ALPHA) OR IFN-ALPHA) (P) PROTEASE
L21
            290 S L21 AND PD<=20020909
L22
            136 DUP REM L22 (154 DUPLICATES REMOVED)
L23
L24
              5 S L23 AND (INTERFERON-ALPHA 2B)
=> D L23 Ibib Abs 108, 116, 123
L23 ANSWER 108 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN
ACCESSION NUMBER:
                         1991:137408 CAPLUS
DOCUMENT NUMBER:
                         114:137408
TITLE:
                         Glycosylated polypeptides for better thermostability
                         and protease resistance
INVENTOR (S):
                         Sasaki, Katsutoshi; Nishi, Tatsunari; Yasumura,
                         Shigeyoshi; Sato, Moriyuki; Itoh, Seiga
                         Kyowa Hakko Kogyo Co., Ltd., Japan
PATENT ASSIGNEE(S):
SOURCE:
                         Eur. Pat. Appl., 130 pp.
                         CODEN: EPXXDW
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                         English
```

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE		
	EP 370205	A2	19900530	EP 1989-117981	19890928 <		
	EP 370205	A 3	19900613				
	EP 370205	B1	19980722				
	R: AT, BE, CH,	DE, ES	, FR, GB, GR	, IT, LI, LU, NL, SE			
	US 5218092	Α	19930608	US 1989-413482	19890927 <		
	JP 02227075	A	19900910	JP 1989-253097	19890928 <		
	JP 2928287	B2	19990803				
	AT 168699	T	19980815	AT 1989-117981	19890928 <		
	ES 2121734	T 3	19981216	ES 1989-117981	19890928 <		
	CA 1341385	С	20020820	CA 1989-614003	19890928 <		
PRIO	RITY APPLN. INFO.:			JP 1988-245705 A	19880929		
AB	Physiol. active poly	/peptide	e-encoding ge	ene is mutagenized such	that		
≥1 new glycosylation sites (markush structure given) are formed.							
The gene is introduced by transformation into, e.g. CHO cells, to produce							
	glycosylated physiol. active polypeptides, e.g. urokinase, containing						
	≥1 new carbohydrate chains. Plasmid pAS28 encoding glycosylated						
	human granulocyte colony stimulating factor hG-CSF[ND28] was constructed						

The gene is introduced by transformation into, e.g. CHO cells, to produce glycosylated physiol. active polypeptides, e.g. urokinase, containing ≥1 new carbohydrate chains. Plasmid pAS28 encoding glycosylated human granulocyte colony stimulating factor hG-CSF[ND28] was constructed and introduced into CHO cells for production The recombinant hG-CSF[ND28] was a mixture of singly and doubly 0-glycosylated froms. The recombinant hG-CSF[ND28] mixture had a better protease-resistance than that of the wild type hG-CSF; and hG-CSF[ND28] having 2 carbohydrate chains had better protease-resistance than that having only one. Glycosylated hG-CSF, hG-CSF[ND28N6], had better thermostability at 56° than the nonglycosylated counterpart obtained by N-glycanase treatment. Glycosylated urokinase, similarly, was prepared. Like natural prourokinase, it scarcely activated the systemic fibrinolytic system; and it had less sensitivity to thrombin and a prolonged plasma elimination half-life (.apprx.2-fold).

L23 ANSWER 116 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN DUPLICATE 57

ACCESSION NUMBER: 1989:590765 CAPLUS

DOCUMENT NUMBER: 111:190765

TITLE: Mass spectrometric analysis of recombinant human

 α -2 interferon

AUTHOR(S): Padron, G.; Besada, V.; Agraz, A.; Quinones, Y.;

Herrera, L.; Shimonishi, Y.; Takao, T.

CORPORATE SOURCE: Cent. Genet. Eng. Biotechnol., Havana, Cuba

SOURCE: Analytica Chimica Acta (1989), 223(2), 361-9

CODEN: ACACAM; ISSN: 0003-2670

DOCUMENT TYPE: Journal LANGUAGE: English

AB Mass spectrometry (MS) was used for the characterization of recombinant human α -2 interferon (α -2 IFN) produced in Escherichia coli. After purification by monoclonal antibody affinity chromatog., α -2 IFN showed two major peaks in reversed-phase liquid chromatog. (RP-LC). Each component was digested with trypsin and Staphylococcus aureus protease V8, sep. or in tandem, and the peptide mixture was analyzed by MS without further purification. The first peak corresponded to the 165 amino acid sequence of human α -2 IFN and the main component of the second peak was the acetylated Cys1 α -2 IFN. It was also possible to verify by MS the location of the S-S bonds in α -2 IFN and the occurrence of incorrect S-S bridges in the products of some renaturation processes. The best renaturation process for obtaining a product without adducts or scrambling of disulfide bonds could be found by using RP-LC and fast-atom-bombardment MS.

L23 ANSWER 123 OF 136 CAPLUS COPYRIGHT 2007 ACS on STN

ACCESSION NUMBER: 1988:144742 CAPLUS

DOCUMENT NUMBER: 108:144742

TITLE: Identification and partial characterization of a novel

protease in Saccharomyces cerevisiae which cleaves the

peptide bond between residues 22 and 23 in α -interferon, and identification of an $\alpha\text{-interferon}$ resistant to said proteolysis

INVENTOR(S): O'Loughlin, John T.

PATENT ASSIGNEE(S):

Interferon Sciences, Inc., USA

SOURCE: Eur. Pat. Appl., 20 pp.

CODEN: EPXXDW

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE		
	EP 240224 EP 240224			EP 1987-302519	19870324 <		
				, IT, LI, LU, NL, SE			
				DK 1987-1614	19870330 <		
	CN 87102497	Α	19871111	CN 1987-102497	19870330 <		
	JP 62296892	Α	19871224	JP 1987-74566	19870330 <		
PRIOF	RITY APPLN. INFO.:			US 1986-845937 A	19860331		
AB	A novel S. cerevisia						
	between basic amino acids at positions 22 and 23, but cleavage does not						
occur if residue 22 is serine. A recombinant interferon .							
	alpha. with serine, threonine, asparagine, glutamine, or glycine						
	at position 22 could be produced intact in a microorganism whose primary						
	proteolytic activity against the natural species is at that site. The						
	protease was partially purified. from a protease						
	-deficient PEP 3-4 S: cerevisiae mutant. It was membrane-bound and						
	activated by the Triton X-100 present during cell lysis. Recombinant						
	interferons α -1, α -2, and α -8 were all						
	incubated with the protease. Both α -2 and α -8 were						
		no acids	s 22 and 23	(which were Arg-Lys and	Arg-Arg,		
		J, ·					

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